

10271-037

(Sheet 1 of 36)

| | |
|---|-----|
| atg agt aat aaa aac gtc aat gta agg aaa tcg cag gaa ata aca ttc Met Ser Asn Lys Asn Val Asn Val Arg Lys Ser Gln Glu Ile Thr Phe 1 5 10 15 | 48 |
| tgc ttg ctg gca ggt atc ctg atg ttc atg gca atg atg gtt gcc gga Cys Leu Leu Ala Gly Ile Leu Met Phe Met Ala Met Met Val Ala Gly 20 25 30 | 96 |
| cgc gct gaa gcg gga gtg gcc tta ggt gcg act cgc gta att tat ccg Arg Ala Glu Ala Gly Val Ala Leu Gly Ala Thr Arg Val Ile Tyr Pro 35 40 45 | 144 |
| gca ggg caa aaa caa gtg caa ctt gcc gtg aca aat aat gat gaa aat Ala Gly Gln Lys Gln Val Gln Leu Ala Val Thr Asn Asn Asp Glu Asn 50 55 60 | 192 |
| agt acc tat tta att caa tca tgg gtg gaa aat gcc gat ggt gta aag Ser Thr Tyr Leu Ile Gln Ser Trp Val Glu Asn Ala Asp Gly Val Lys 65 70 75 80 | 240 |
| gat ggt cgt ttt atc gtg acg cct cct ctg ttt gcg atg aag gga aaa Asp Gly Arg Phe Ile Val Thr Pro Pro Leu Phe Ala Met Lys Gly Lys 85 90 95 | 288 |
| aaa gag aat acc tta cgt att ctt gat gca aca aat aac caa ttg cca Lys Glu Asn Thr Leu Arg Ile Leu Asp Ala Thr Asn Asn Gln Leu Pro 100 105 110 | 336 |
| cag gac cgg gaa agt tta ttc tgg atg aac gtt aaa gcg att ccg tca Gln Asp Arg Glu Ser Leu Phe Trp Met Asn Val Lys Ala Ile Pro Ser 115 120 125 | 384 |
| atg gat aaa tca aaa ttg act gag aat acg cta cag ctc gca att atc Met Asp Lys Ser Lys Leu Thr Glu Asn Thr Leu Gln Leu Ala Ile Ile 130 135 140 | 432 |
| agc cgc att aaa ctg tac tat cgc ccg gct aaa tta gcg ttg cca ccc Ser Arg Ile Lys Leu Tyr Tyr Arg Pro Ala Lys Leu Ala Leu Pro P o 145 150 155 160 | 480 |
| gat cag gcc gca gaa aaa tta aga ttt cgt cgt agc gcg aat tct cgt Asp Gln Ala Ala Glu Lys Leu Arg Phe Arg Arg Ser Ala Asn Ser Leu 165 170 175 | 528 |
| acg ctg att aac ccg aca ccc tat tac ctg acg gta aca gag ttg aat Thr Leu Ile Asn Pro Thr Pro Tyr Tyr Leu Thr Val Thr Glu Leu Asn 180 185 190 | 576 |
| gcc gga acc cgg gtt ctt gaa aat gca ttg gtg cct cca atg ggc gaa Ala Gly Thr Arg Val Leu Glu Asn Ala Leu Val Pro Pro Met Gly G u 195 200 205 | 624 |
| agc acg gtt aaa ttg cct tct gat gca gga agc aat att act tac cga Ser Thr Val Lys Leu Pro Ser Asp Ala Gly Ser Asn Ile Thr Tyr Arg 210 215 220 | 672 |
| aca ata aat gat tat ggc gca ctt acc ccc aaa atg acg ggc gta aag Thr Ile Asn Asp Tyr Gly Ala Leu Thr Pro Lys Met Thr Gly Val Met 225 230 235 240 | 720 |
| gaa taa Glu | 726 |

Figure 1A

Met Ser Asn Lys Asn Val Asn Val Arg Lys Ser Gln Glu Ile Thr Phe
 1 5 10 15
 Cys Leu Leu Ala Gly Ile Leu Met Phe Met Ala Met Met Val Ala Gly
 20 25 30
 Arg Ala Glu Ala Gly Val Ala Leu Gly Ala Thr Arg Val Ile Tyr Pro
 35 40 45
 Ala Gly Gln Lys Gln Val Gln Leu Ala Val Thr Asn Asn Asp Glu Asn
 50 55 60
 Ser Thr Tyr Leu Ile Gln Ser Trp Val Glu Asn Ala Asp Gly Val Lys
 65 70 75 80
 Asp Gly Arg Phe Ile Val Thr Pro Pro Leu Phe Ala Met Lys Gly Lys
 85 90 95
 Lys Glu Asn Thr Leu Arg Ile Leu Asp Ala Thr Asn Asn Gln Leu Pro
 100 105 110
 Gln Asp Arg Glu Ser Leu Phe Trp Met Asn Val Lys Ala Ile Pro Ser
 115 120 125
 Met Asp Lys Ser Lys Leu Thr Glu Asn Thr Leu Gln Leu Ala Ile Ile
 130 135 140
 Ser Arg Ile Lys Leu Tyr Tyr Arg Pro Ala Lys Leu Ala Leu Pro Pro
 145 150 155 160
 Asp Gln Ala Ala Glu Lys Leu Arg Phe Arg Arg Ser Ala Asn Ser Leu
 165 170 175
 Thr Leu Ile Asn Pro Thr Pro Tyr Tyr Leu Thr Val Thr Glu Leu Asn
 180 185 190
 Ala Gly Thr Arg Val Leu Glu Asn Ala Leu Val Pro Pro Met Gly Glu
 195 200 205
 Ser Thr Val Lys Leu Pro Ser Asp Ala Gly Ser Asn Ile Thr Tyr Arg
 210 215 220
 Thr Ile Asn Asp Tyr Gly Ala Leu Thr Pro Lys Met Thr Gly Val Met
 225 230 235 240
 Glu

Figure 1B

| | | | | | | | | | | | | | | | | |
|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|
| atg | aaa | cga | ggt | att | acc | ctg | ttt | gct | gta | ctg | ctg | atg | ggc | tgg | tgg | 48 |
| Met | Lys | Arg | Val | Ile | Thr | Leu | Phe | Ala | Val | Leu | Leu | Met | Gly | Trp | Ser | |
| | -20 | | | | | -15 | | | | | -10 | | | | | |
| gta | aat | gcc | tgg | tca | ttc | gcc | tgt | aaa | acc | gcc | aat | ggt | acc | gct | atc | 96 |
| Val | Asn | Ala | Trp | Ser | Phe | Ala | Cys | Lys | Thr | Ala | Asn | Gly | Thr | Ala | Ile | |
| -5 | | | | -1 | 1 | | | | 5 | | | | | 10 | | |
| cct | att | ggc | ggt | ggc | agc | gcc | aat | ggt | tat | gta | aac | ctt | gcg | ccc | gtc | 144 |
| Pro | Ile | Gly | Gly | Gly | Ser | Ala | Asn | Val | Tyr | Val | Asn | Leu | Ala | Pro | Val | |
| | | 15 | | | | | | 20 | | | | | 25 | | | |
| gtg | aat | gtg | ggg | caa | aac | ctg | gtc | gtg | gat | ctt | tgc | acg | caa | atc | ttt | 192 |
| Val | Asn | Val | Gly | Gln | Asn | Leu | Val | Val | Asp | Leu | Ser | Thr | Gln | Ile | Phe | |
| | 30 | | | | | | 35 | | | | | 40 | | | | |
| tgc | cat | aac | gat | tat | ccg | gaa | acc | att | aca | gac | tat | gtc | aca | ctg | caa | 240 |
| Cys | His | Asn | Asp | Tyr | Pro | Glu | Thr | Ile | Thr | Asp | Tyr | Val | Thr | Leu | Gln | |
| | 45 | | | | | 50 | | | | | 55 | | | | | |
| cga | ggc | tgc | gct | tat | ggc | ggc | gtg | tta | tct | aat | ttt | tcc | ggg | acc | gta | 288 |
| Arg | Gly | Ser | Ala | Tyr | Gly | Gly | Val | Leu | Ser | Asn | Phe | Ser | Gly | Thr | Val | |
| 60 | | | | | 65 | | | | 70 | | | | | | 75 | |
| aaa | tat | agt | ggc | agt | agc | tat | cca | ttt | cct | acc | acc | agc | gaa | acg | ccg | 336 |
| Lys | Tyr | Ser | Gly | Ser | Ser | Tyr | Pro | Phe | Pro | Thr | Thr | Ser | Glu | Thr | Pro | |
| | | | | 80 | | | | | 85 | | | | | 90 | | |
| cgc | ggt | ggt | tat | aat | tgc | aga | acg | gat | aag | ccg | tgg | ccg | gtg | gcg | ctt | 384 |
| Arg | Val | Val | Tyr | Asn | Ser | Arg | Thr | Asp | Lys | Pro | Trp | Pro | Val | Ala | Leu | |
| | | | 95 | | | | | 100 | | | | | 105 | | | |
| tat | ttg | acg | cct | gtg | agc | agt | gcg | ggc | ggg | gtg | gcg | att | aaa | gct | ggc | 432 |
| Tyr | Leu | Thr | Pro | Val | Ser | Ser | Ala | Gly | Gly | Val | Ala | Ile | Lys | Ala | Gly | |
| | 110 | | | | | | 115 | | | | | 120 | | | | |
| tca | tta | att | gcc | gtg | ctt | att | ttg | cga | cag | acc | aac | aac | tat | aac | agc | 480 |
| Ser | Leu | Ile | Ala | Val | Leu | Ile | Leu | Arg | Gln | Thr | Asn | Asn | Tyr | Asn | Ser | |
| | 125 | | | | | 130 | | | | | 135 | | | | | |
| gat | gat | ttc | cag | ttt | gtg | tgg | aat | att | tac | gcc | aat | aat | gat | gtg | gtg | 528 |
| Asp | Asp | Phe | Gln | Phe | Val | Trp | Asn | Ile | Tyr | Ala | Asn | Asn | Asp | Val | Val | |
| 140 | | | | | 145 | | | | | 150 | | | | | 155 | |
| gtg | cct | act | ggc | ggc | tgc | gat | ggt | tct | gct | cgt | gat | gtc | acc | ggt | act | 576 |
| Val | Pro | Thr | Gly | Gly | Cys | Asp | Val | Ser | Ala | Arg | Asp | Val | Thr | Val | Thr | |
| | | | 160 | | | | | | 165 | | | | | 170 | | |
| ctg | ccg | gac | tac | cct | ggg | tca | gtg | cca | att | cct | ctt | acc | ggt | tat | tgt | 624 |
| Leu | Pro | Asp | Tyr | Pro | Gly | Ser | Val | Pro | Ile | Pro | Leu | Thr | Val | Tyr | Cys | |
| | | | 175 | | | | | 180 | | | | | | 185 | | |

Figure 1C

| | |
|---|-----|
| gcg aaa agc caa aac ctg ggg tat tac ctc tcc ggc aca acc gca cat | 672 |
| Ala Lys Ser Gln Asn Leu Gly Tyr Tyr Leu Ser Gly Thr Thr Ala Asp | |
| 190 195 200 | |
| gcg ggc aac tcg att ttc acc aat acc gcg tcg ttt tca cct gca cag | 720 |
| Ala Gly Asn Ser Ile Phe Thr Asn Thr Ala Ser Phe Ser Pro Ala Gln | |
| 205 210 215 | |
| ggc gtc ggc gta cag ttg acg cgc aac ggt acg att att cca gcg aat | 768 |
| Gly Val Gly Val Gln Leu Thr Arg Asn Gly Thr Ile Ile Pro Ala Asn | |
| 220 225 230 235 | |
| aac acg gta tcg tta gga gca gta ggg act tcg gcg gtg agt ctg gga | 816 |
| Asn Thr Val Ser Leu Gly Ala Val Gly Thr Ser Ala Val Ser Leu Gly | |
| 240 245 250 | |
| tta acg gca aat tat gca cgt acc gga ggg cag gtg act gca ggg aat | 864 |
| Leu Thr Ala Asn Tyr Ala Arg Thr Gly Gly Gln Val Thr Ala Gly Asn | |
| 255 260 265 | |
| gtg caa tcg att att ggc gtg act ttt gtt tat caa taa | 903 |
| Val Gln Ser Ile Ile Gly Val Thr Phe Val Tyr Gln | |
| 270 275 | |

Figure 1C (Cont'd)

Met Lys Arg Val Ile Thr Leu Phe Ala Val Leu Leu Met Gly Trp Ser
 -20 -15 -10
 Val Asn Ala Trp Ser Phe Ala Cys Lys Thr Ala Asn Gly Thr Ala Ile
 -5 -1 1 5 10
 Pro Ile Gly Gly Gly Ser Ala Asn Val Tyr Val Asn Leu Ala Pro Val
 15 20 25
 Val Asn Val Gly Gln Asn Leu Val Val Asp Leu Ser Thr Gln Ile Phe
 30 35 40
 Cys His Asn Asp Tyr Pro Glu Thr Ile Thr Asp Tyr Val Thr Leu Gln
 45 50 55
 Arg Gly Ser Ala Tyr Gly Gly Val Leu Ser Asn Phe Ser Gly Thr Val
 60 65 70 75
 Lys Tyr Ser Gly Ser Ser Tyr Pro Phe Pro Thr Thr Ser Glu Thr Pro
 80 85 90
 Arg Val Val Tyr Asn Ser Arg Thr Asp Lys Pro Trp Pro Val Ala Leu
 95 100 105
 Tyr Leu Thr Pro Val Ser Ser Ala Gly Gly Val Ala Ile Lys Ala Gly
 110 115 120
 Ser Leu Ile Ala Val Leu Ile Leu Arg Gln Thr Asn Asn Tyr Asn Ser
 125 130 135
 Asp Asp Phe Gln Phe Val Trp Asn Ile Tyr Ala Asn Asn Asp Val Val
 140 145 150 155
 Val Pro Thr Gly Gly Cys Asp Val Ser Ala Arg Asp Val Thr Val Thr
 160 165 170
 Leu Pro Asp Tyr Pro Gly Ser Val Pro Ile Pro Leu Thr Val Tyr Cys
 175 180 185
 Ala Lys Ser Gln Asn Leu Gly Tyr Tyr Leu Ser Gly Thr Thr Ala Asp
 190 195 200
 Ala Gly Asn Ser Ile Phe Thr Asn Thr Ala Ser Phe Ser Pro Ala Gln
 205 210 215
 Gly Val Gly Val Gln Leu Thr Arg Asn Gly Thr Ile Ile Pro Ala Asn
 220 225 230 235
 Asn Thr Val Ser Leu Gly Ala Val Gly Thr Ser Ala Val Ser Leu Gly
 240 245 250
 Leu Thr Ala Asn Tyr Ala Arg Thr Gly Gly Gln Val Thr Ala Gly Asn
 255 260 265
 Val Gln Ser Ile Ile Gly Val Thr Phe Val Tyr Gln
 270 275

Figure 1D

a



b

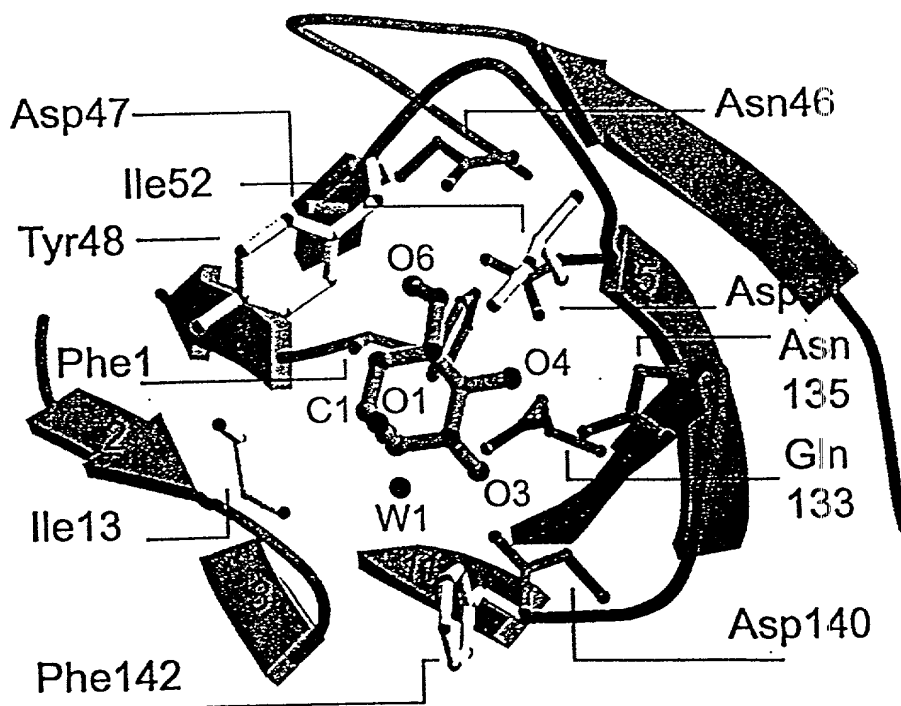
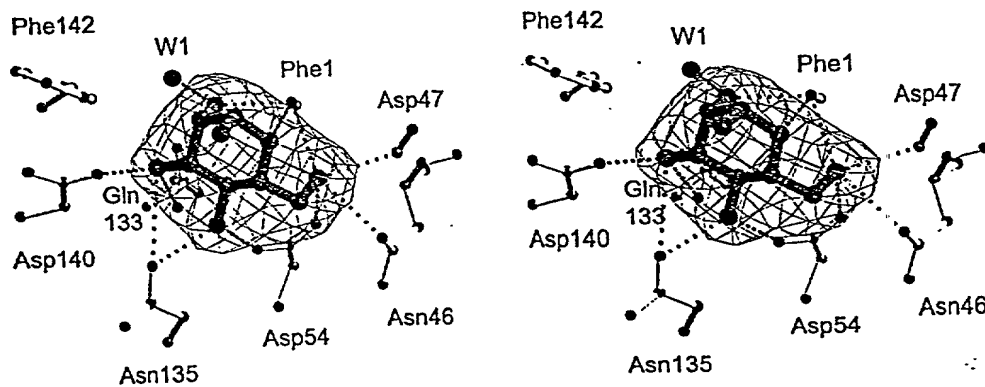


Figure 2
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FOOT2F.S805FOOT

C

10271-037



d

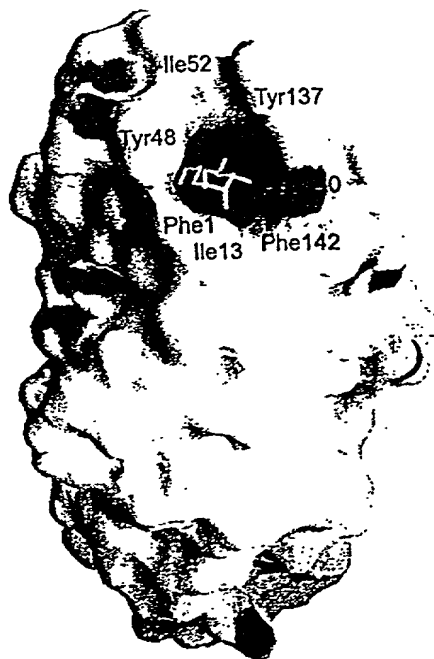


Figure 2
Sheet 7 of 36

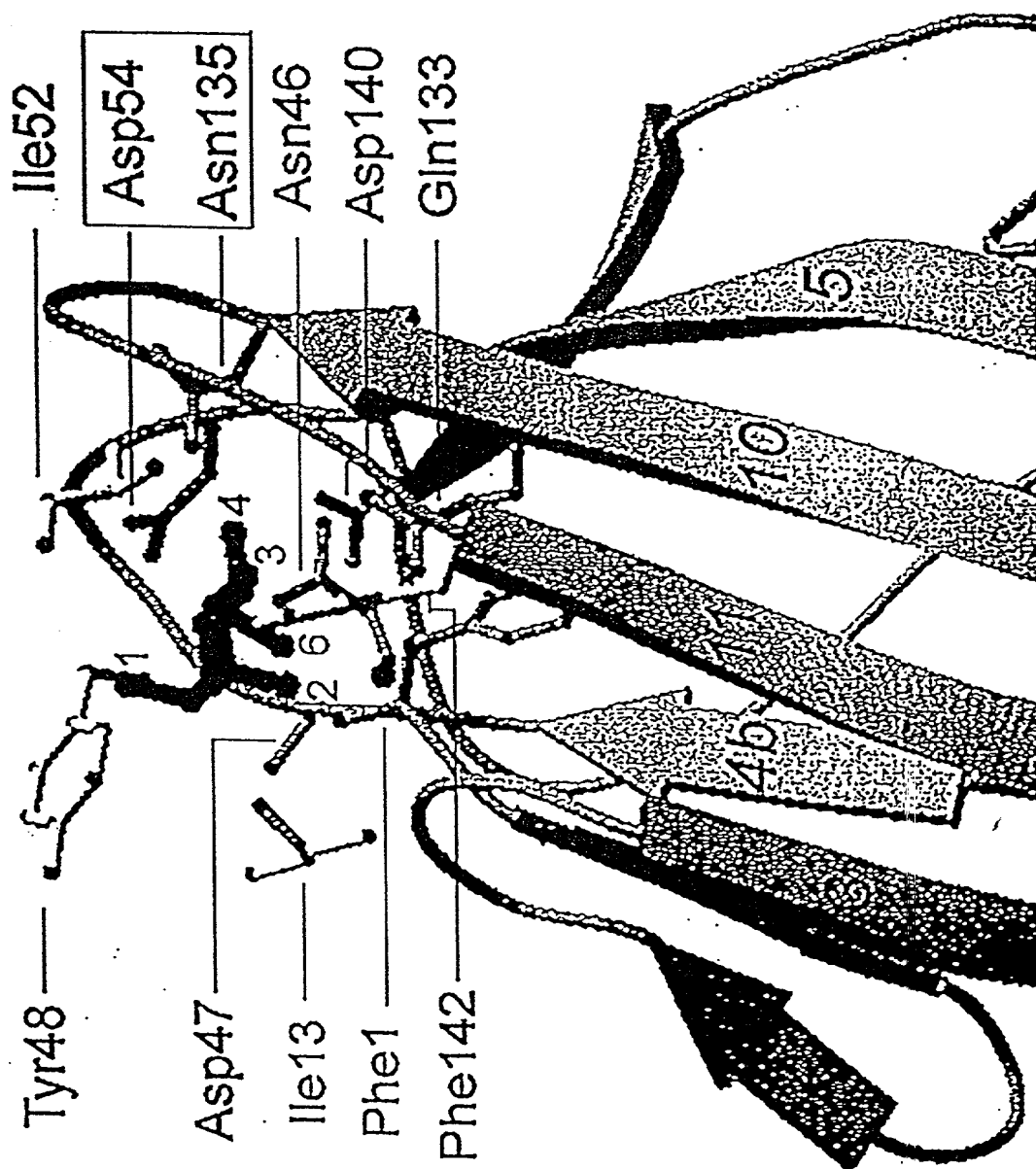


Figure 2
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Figure 3
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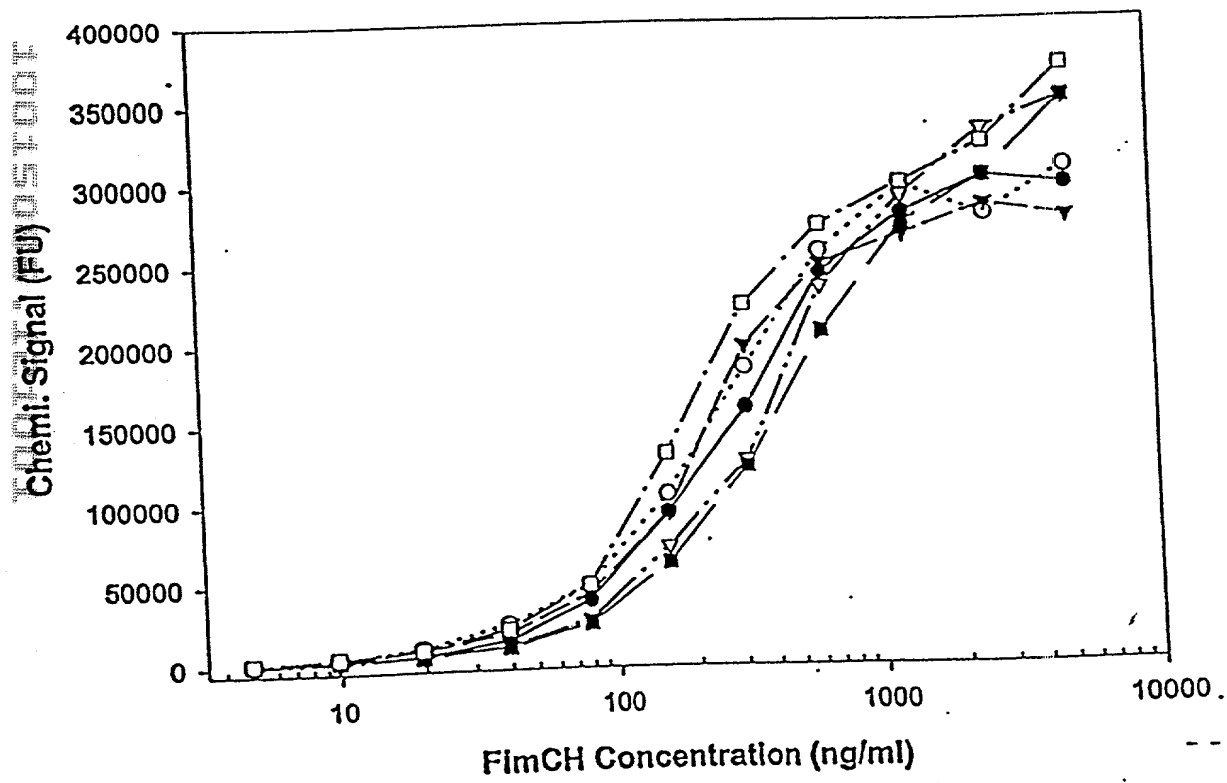


Figure 4
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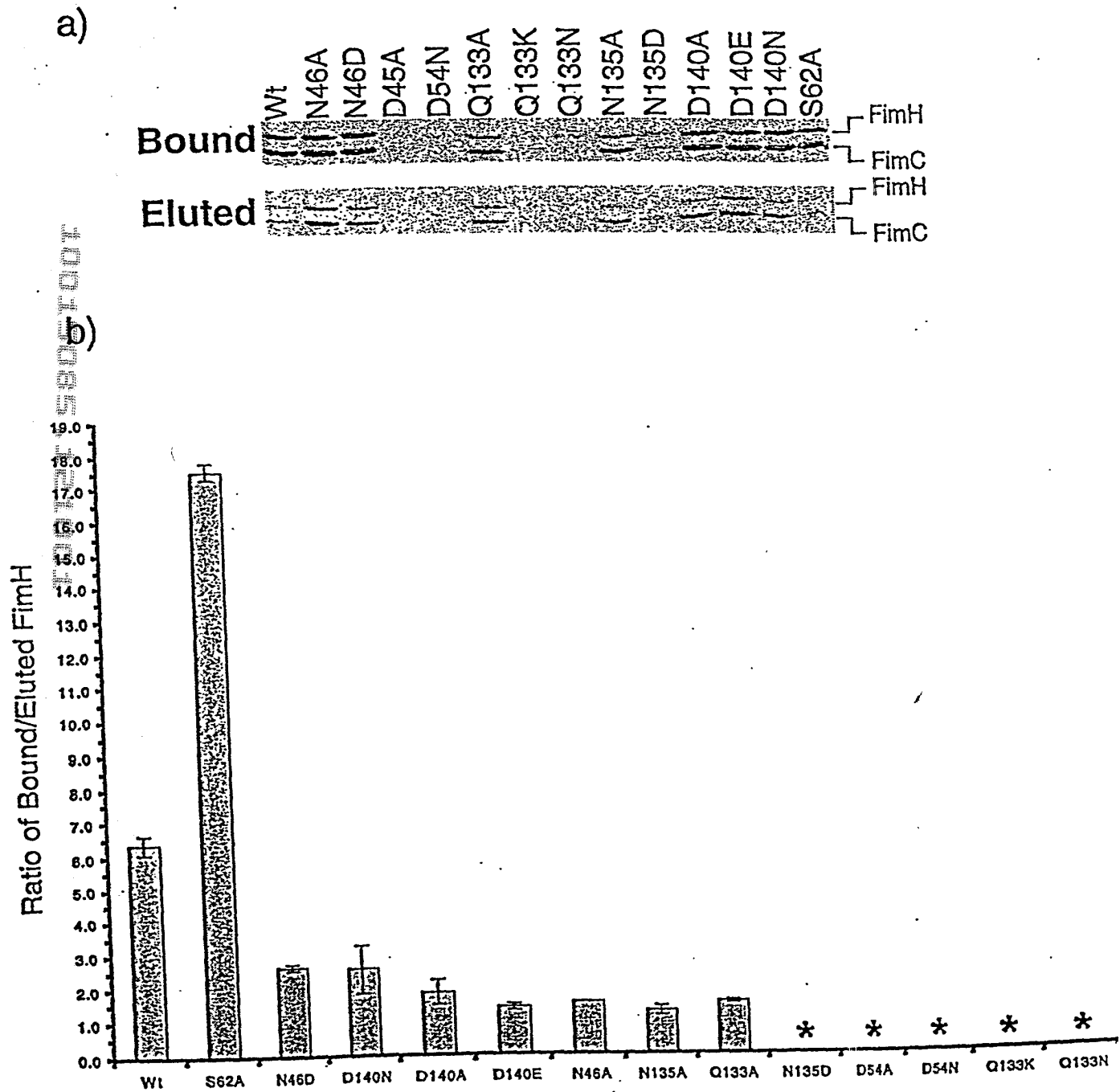


Figure 5
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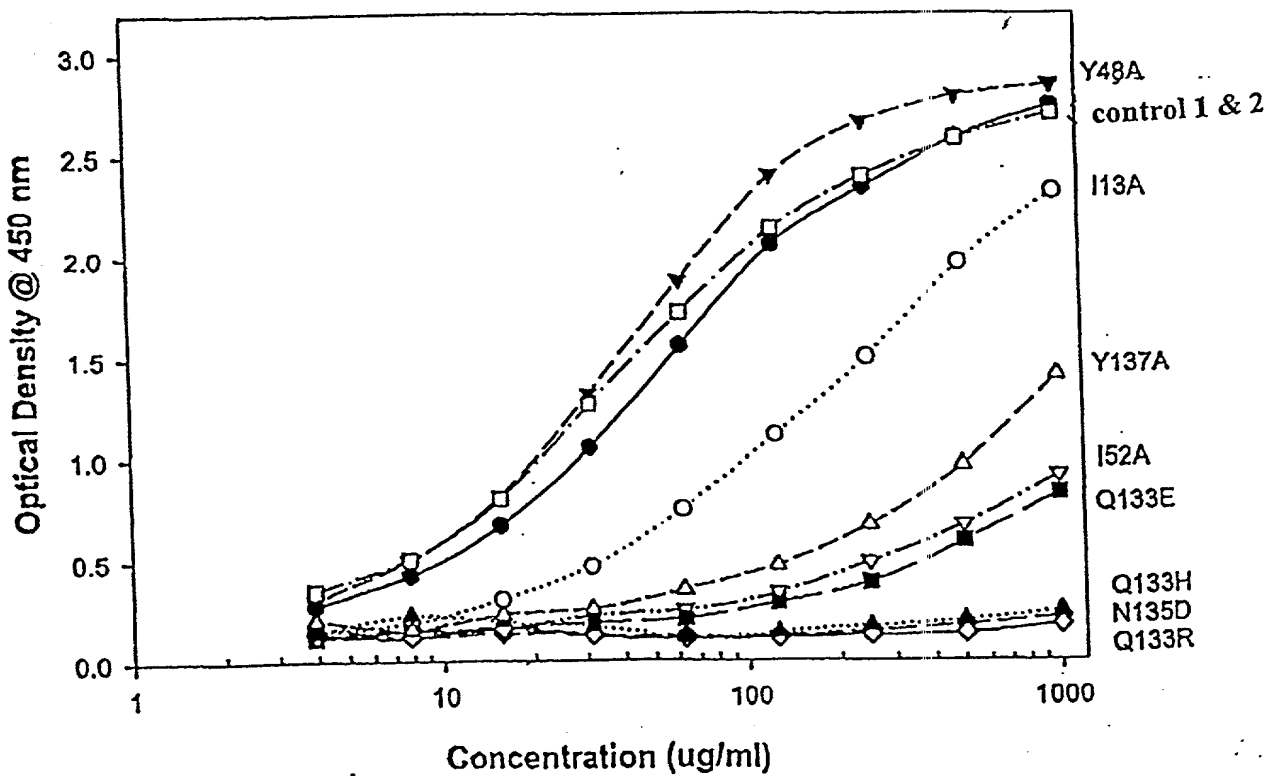
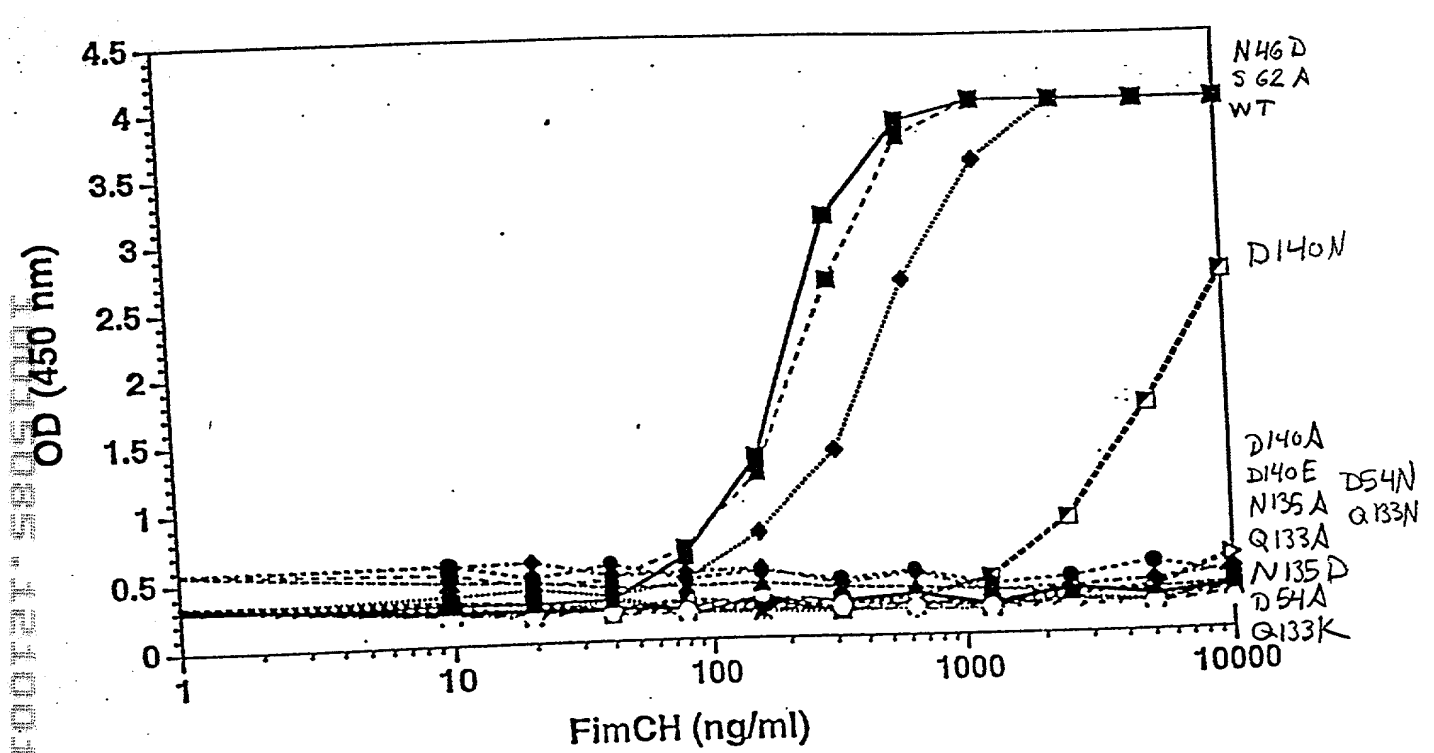


Figure 6
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B)

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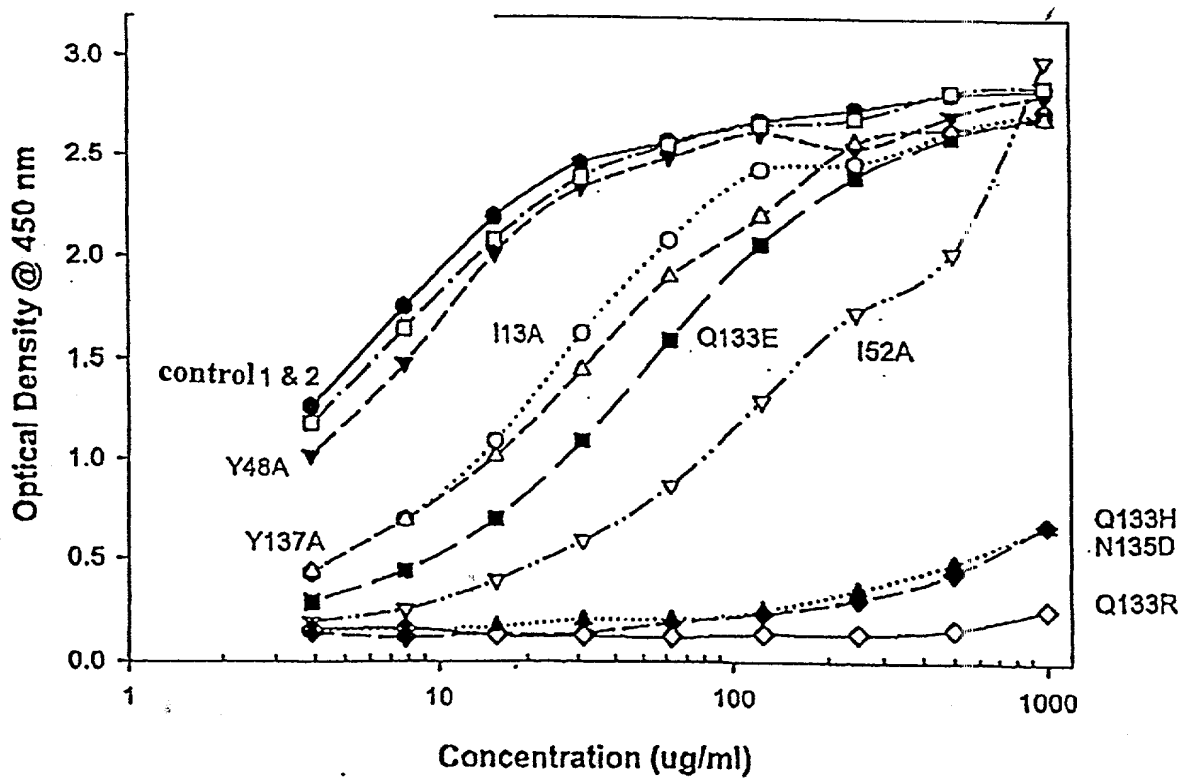
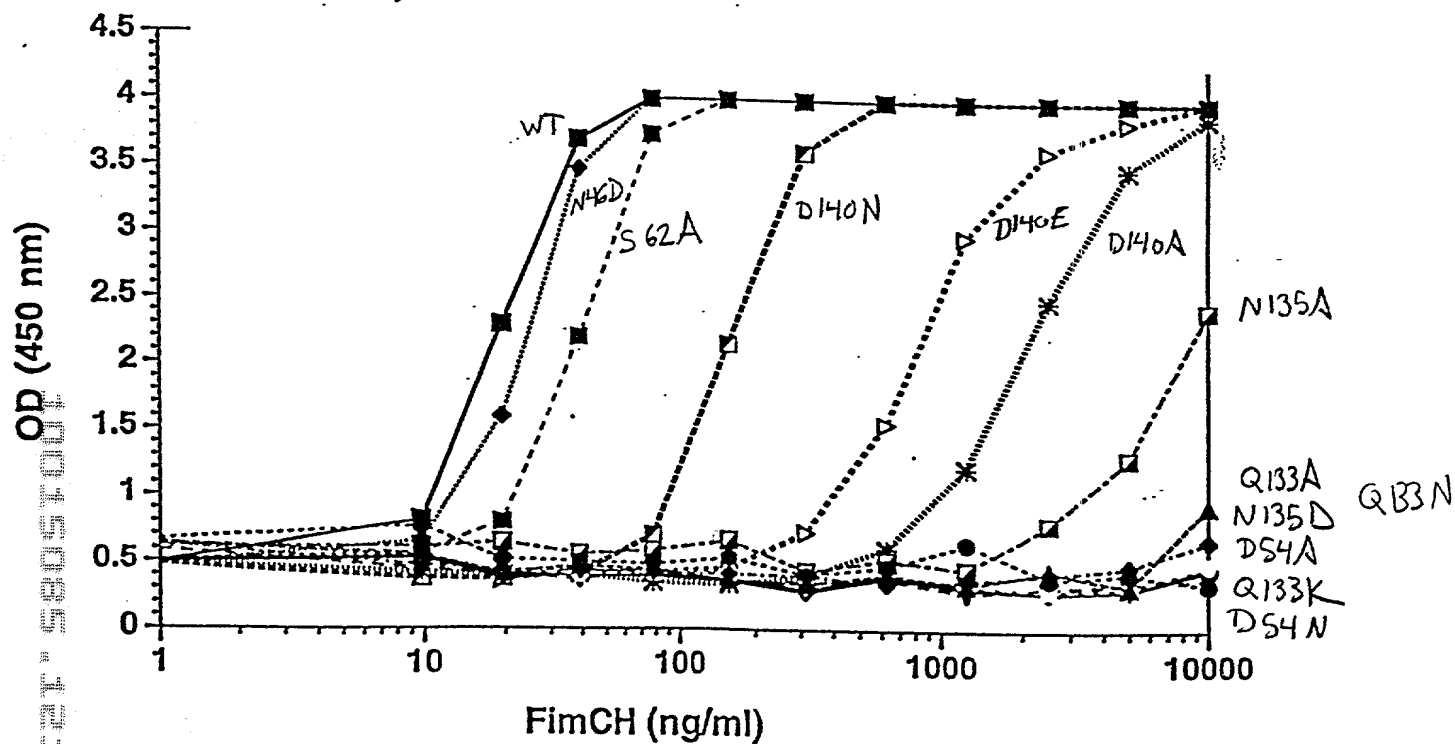
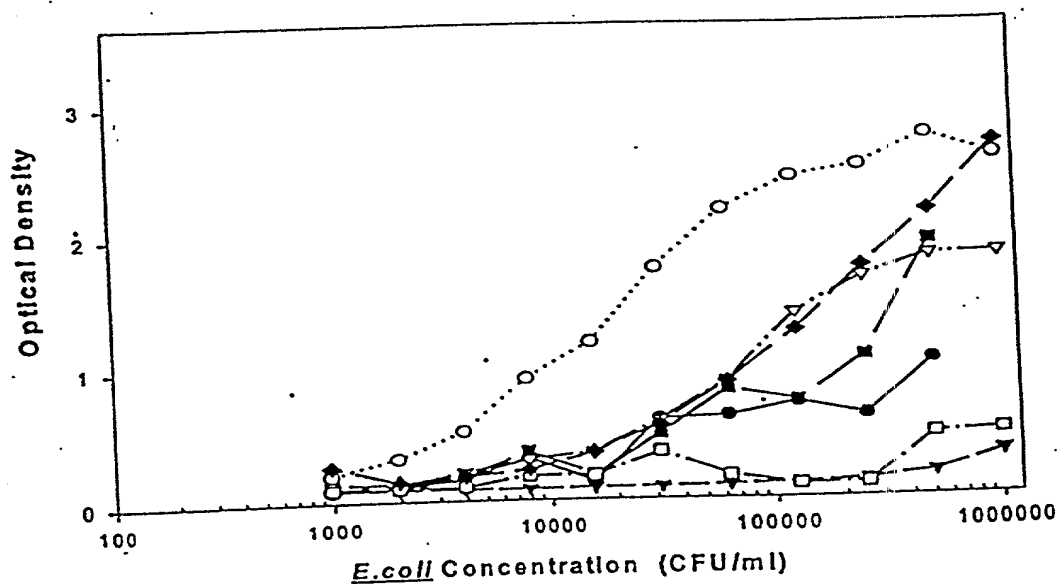


Figure 6

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A



B

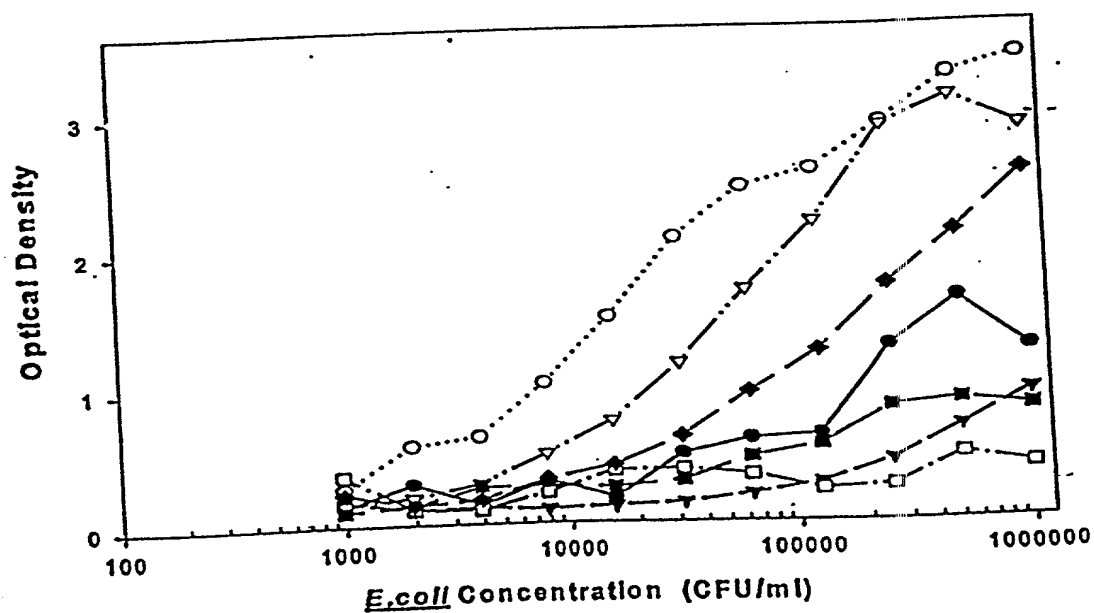


Figure 7
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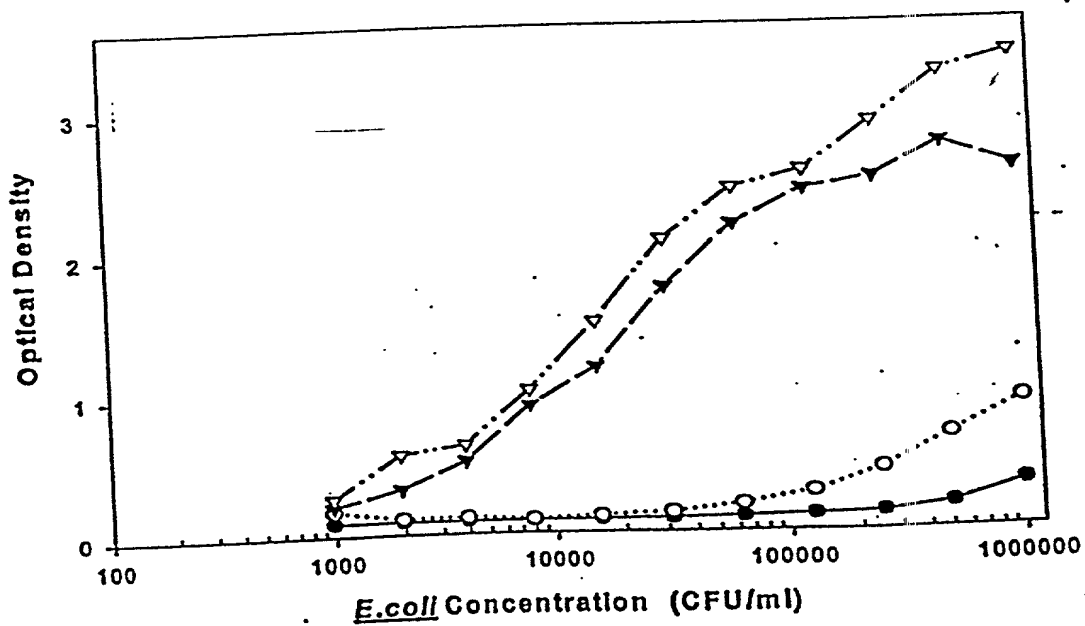
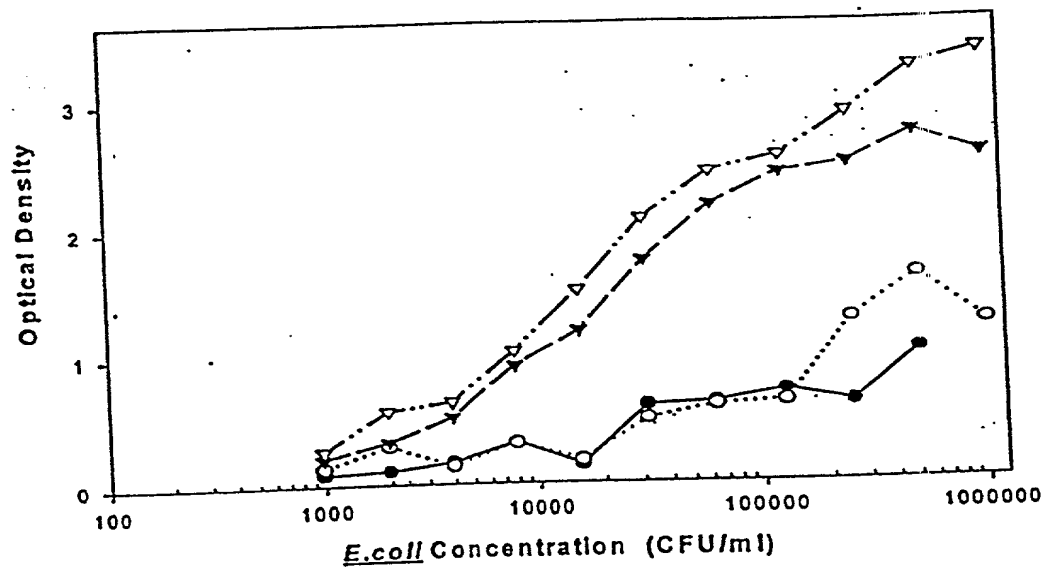


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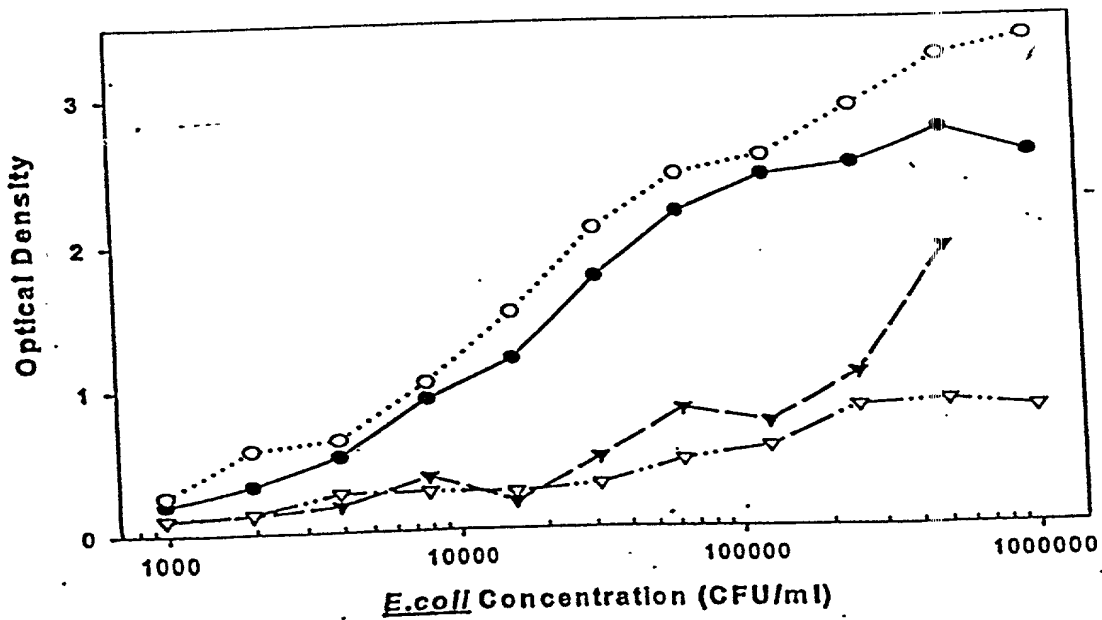
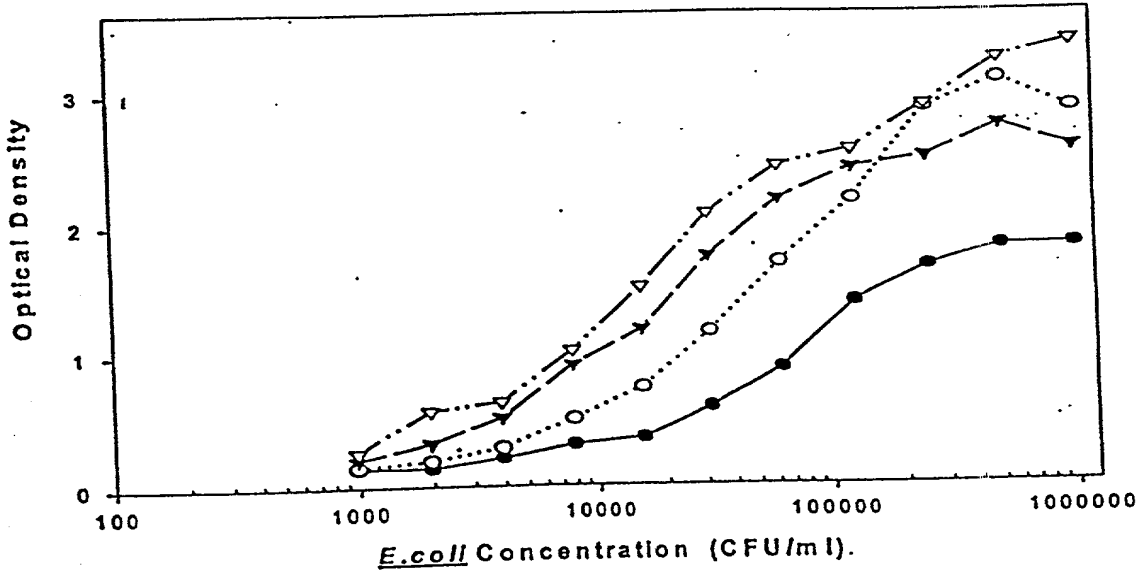


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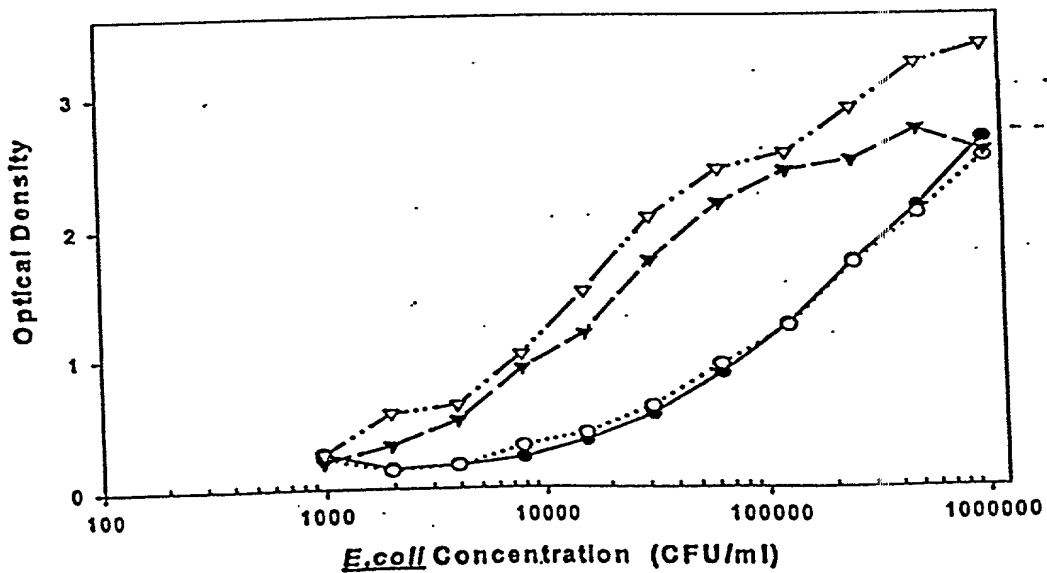
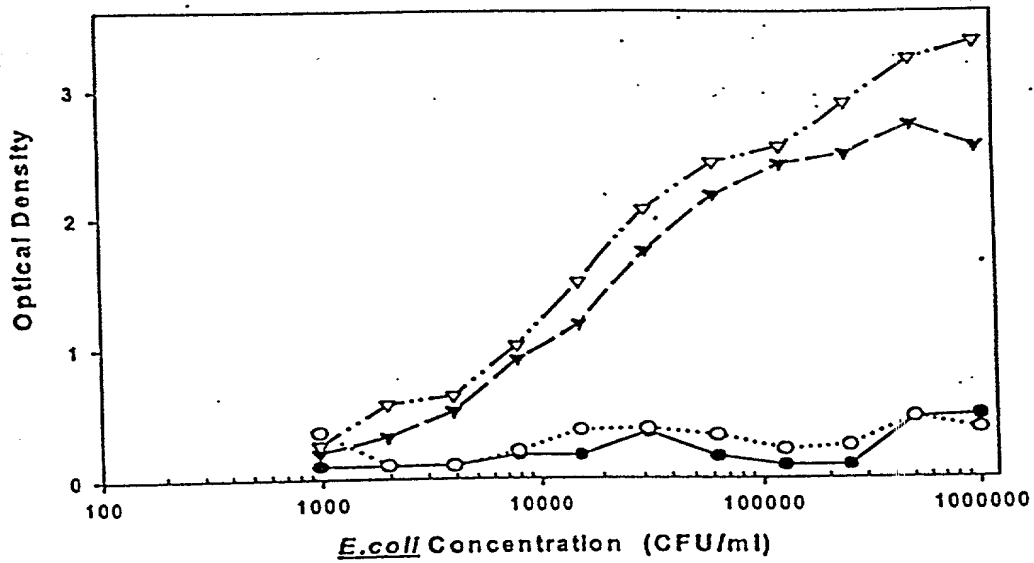


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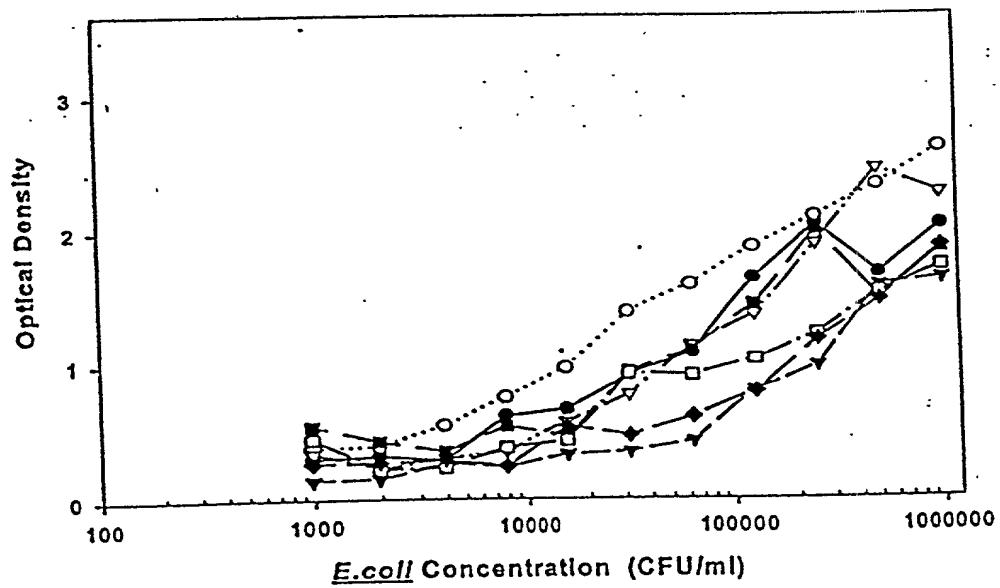
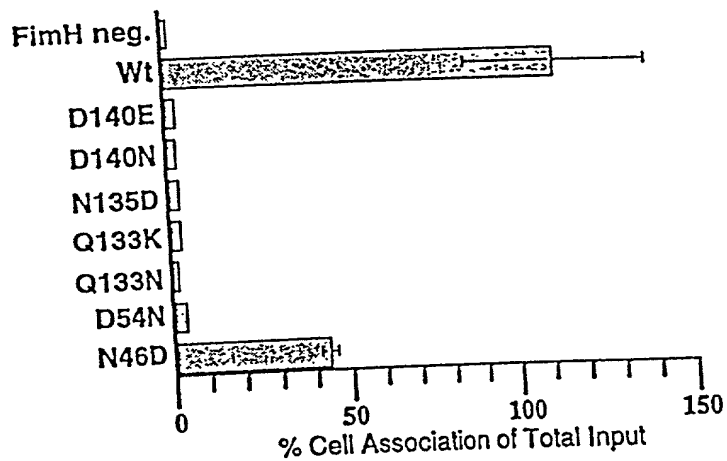
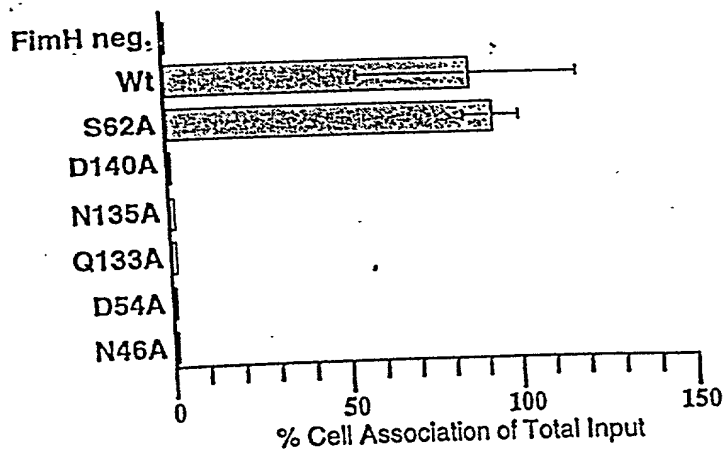


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a)



B)

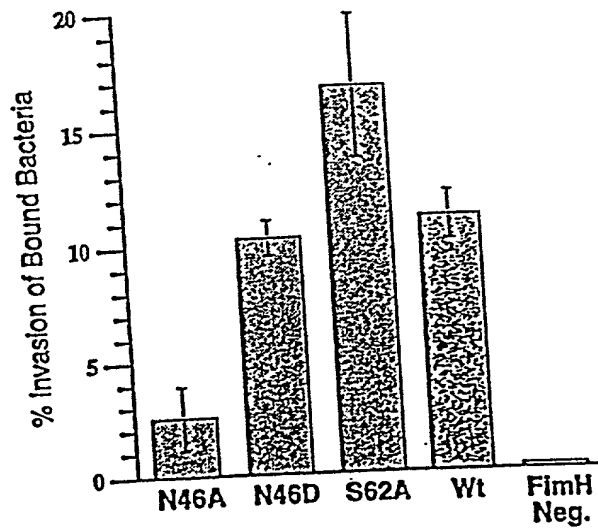


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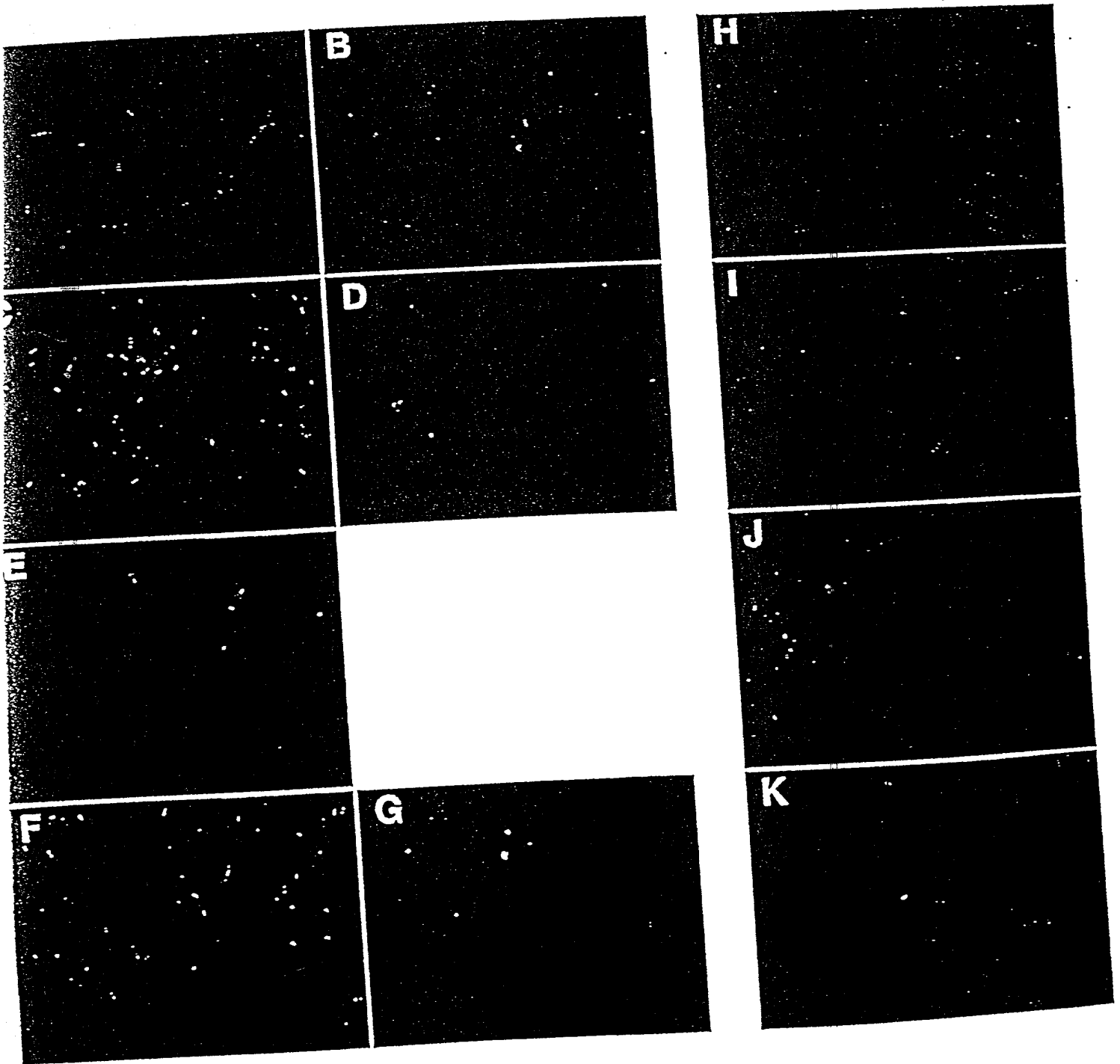
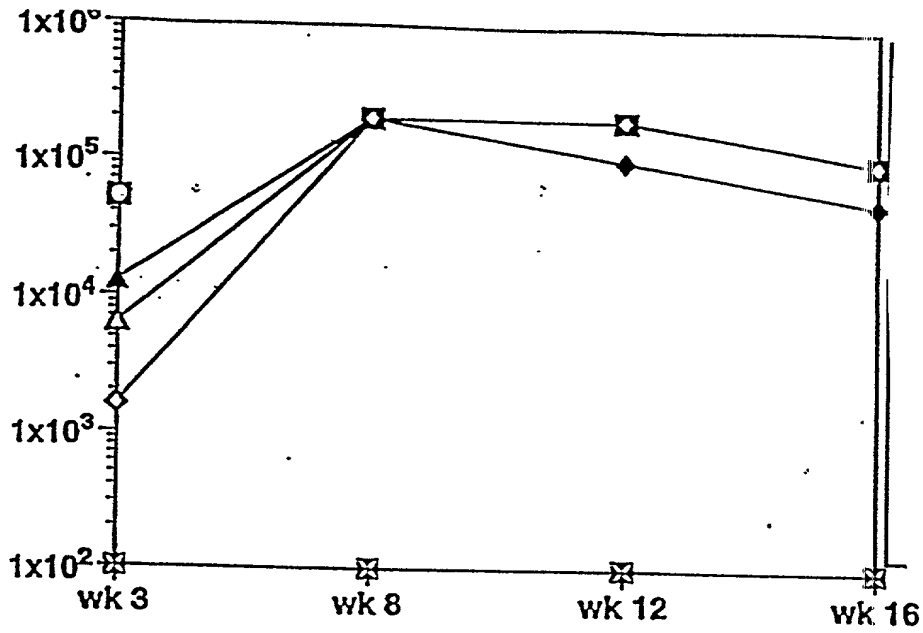
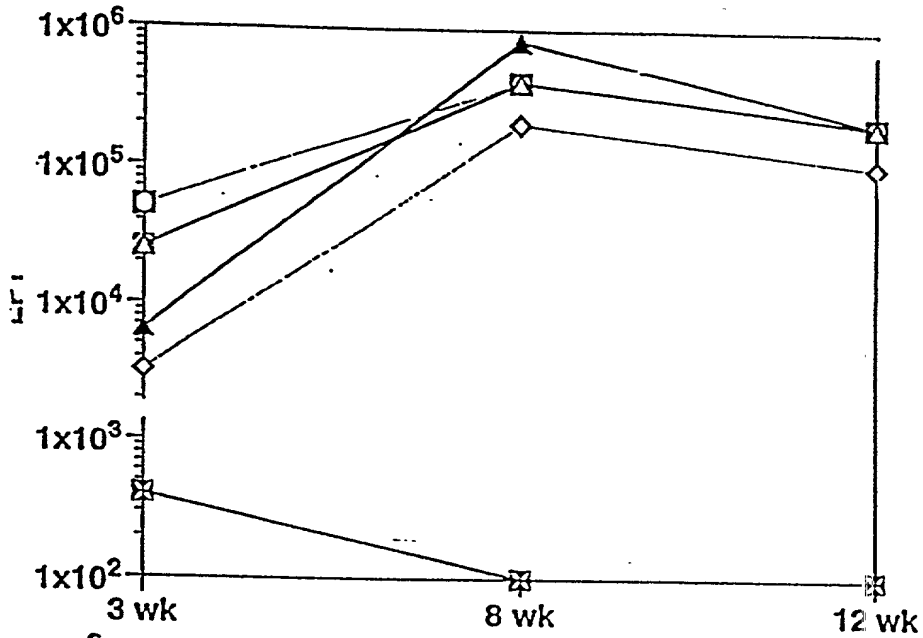


Figure 9
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A



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C

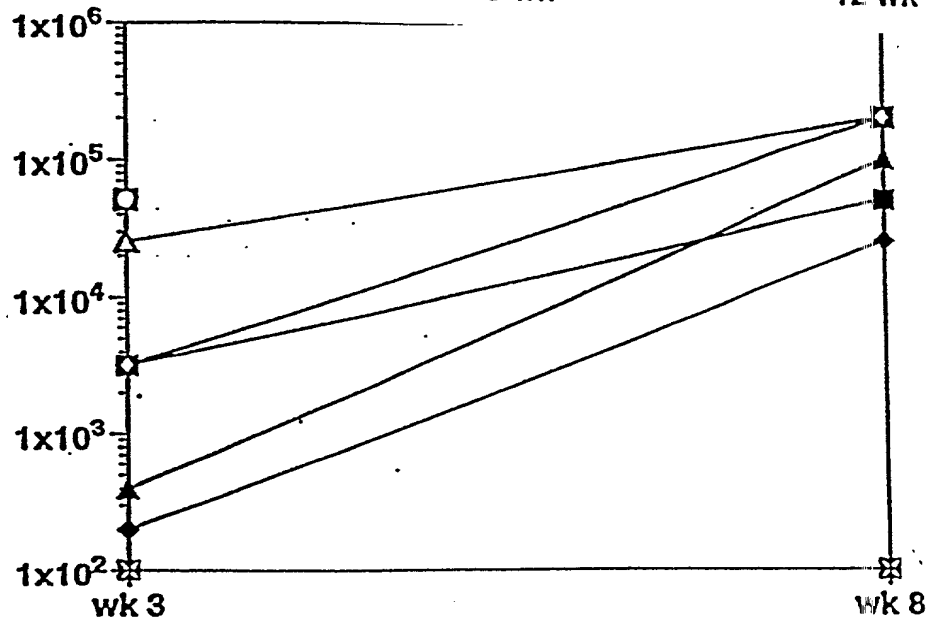
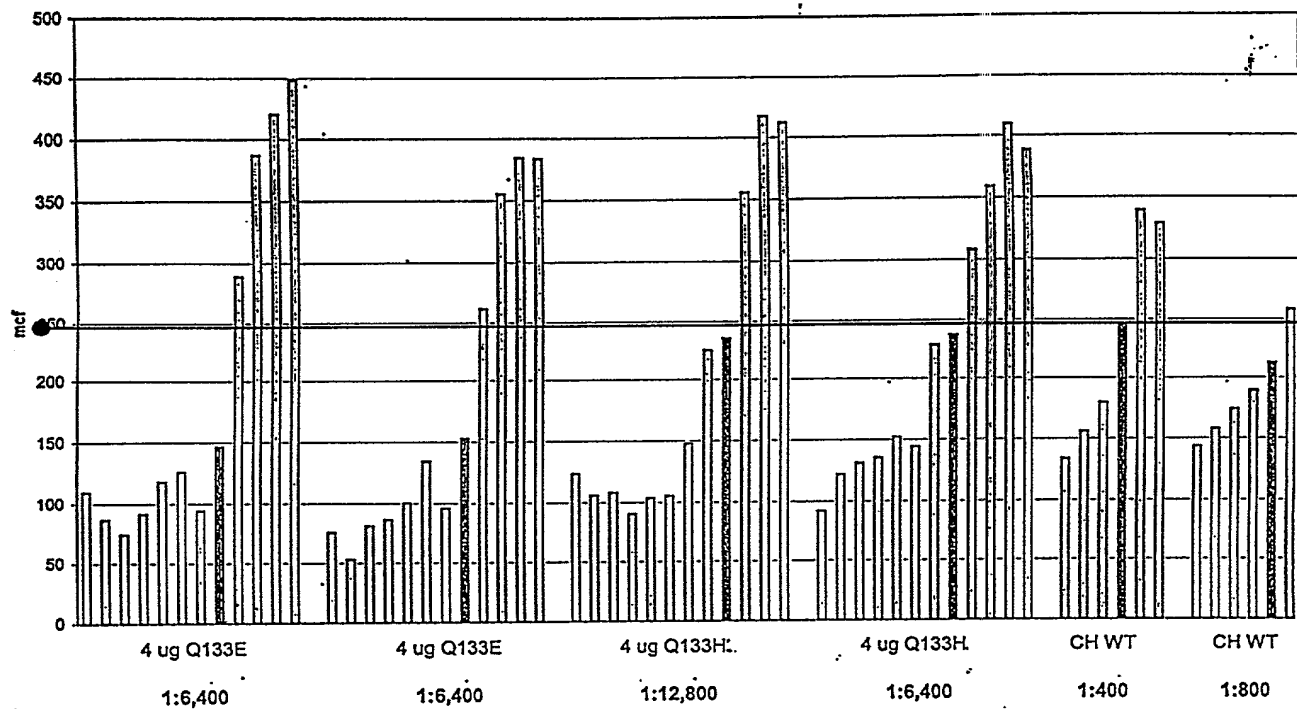


Figure 10
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A



B

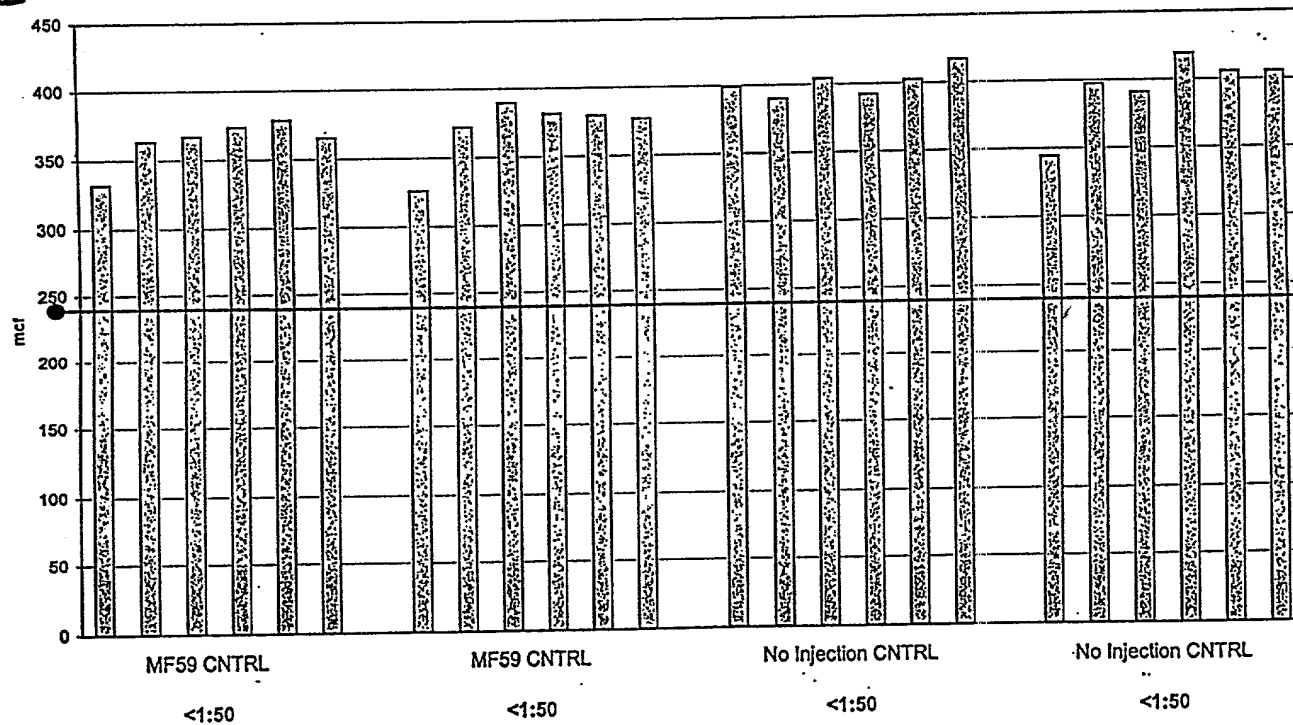


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C

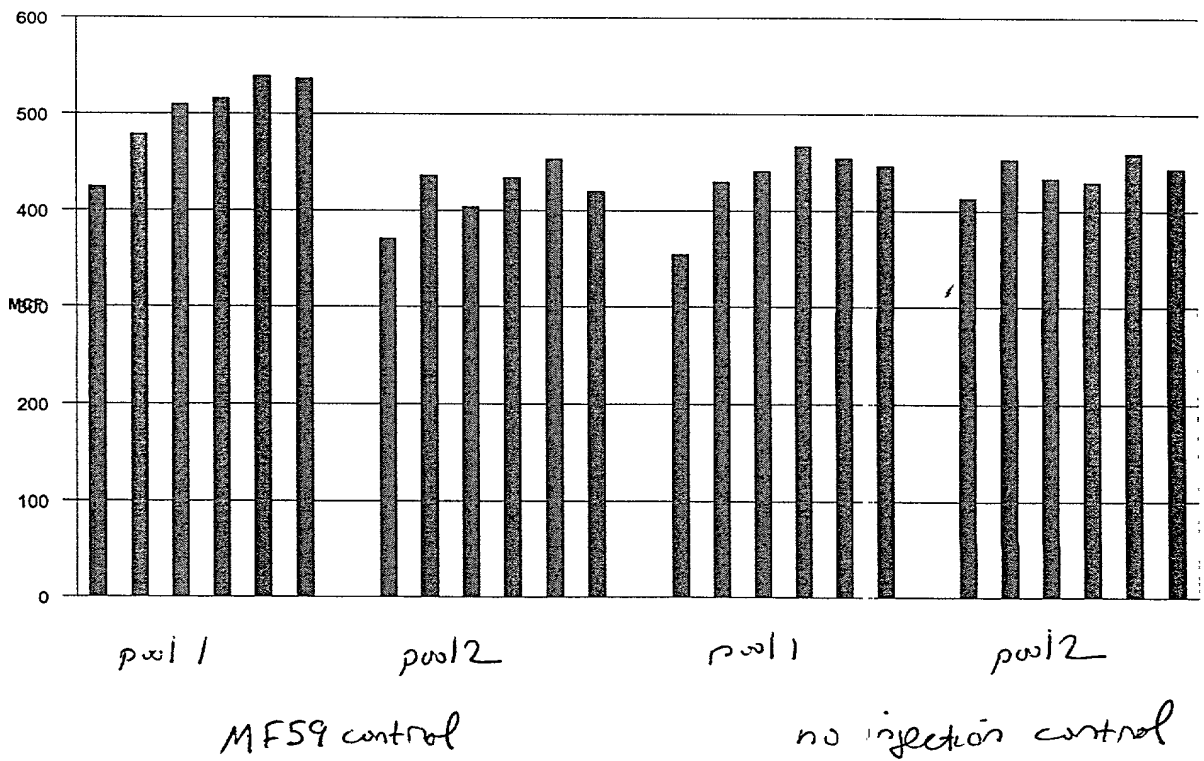
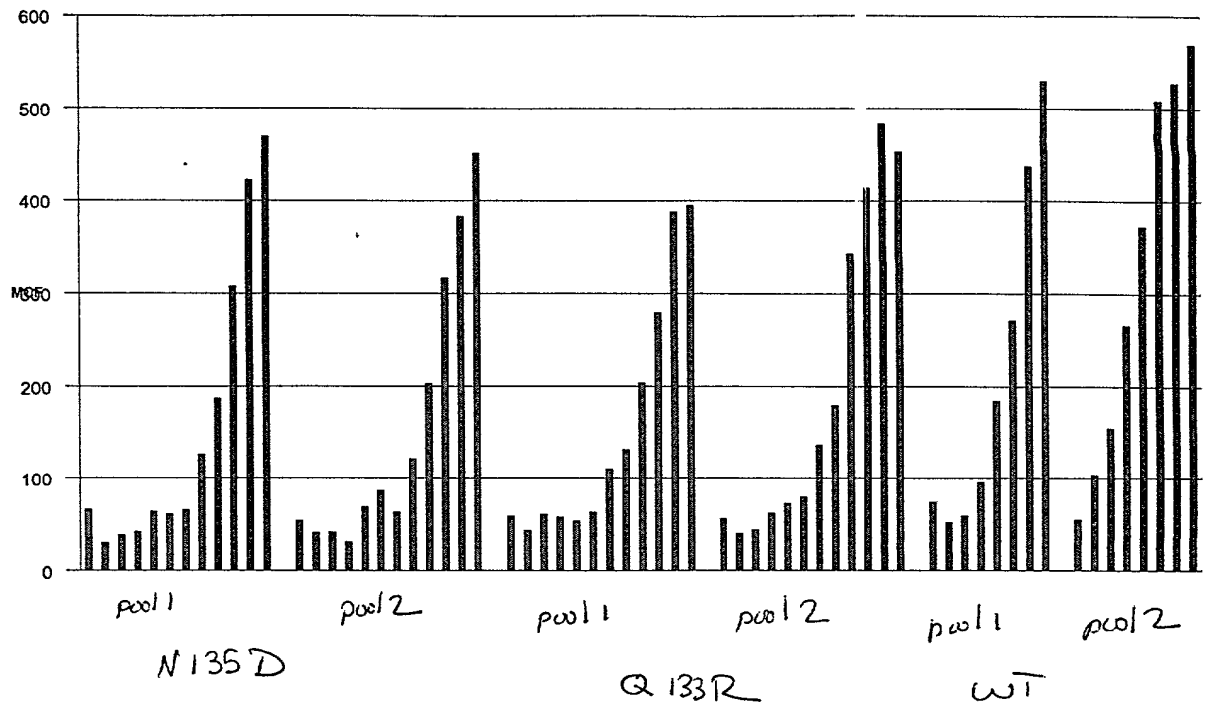


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A

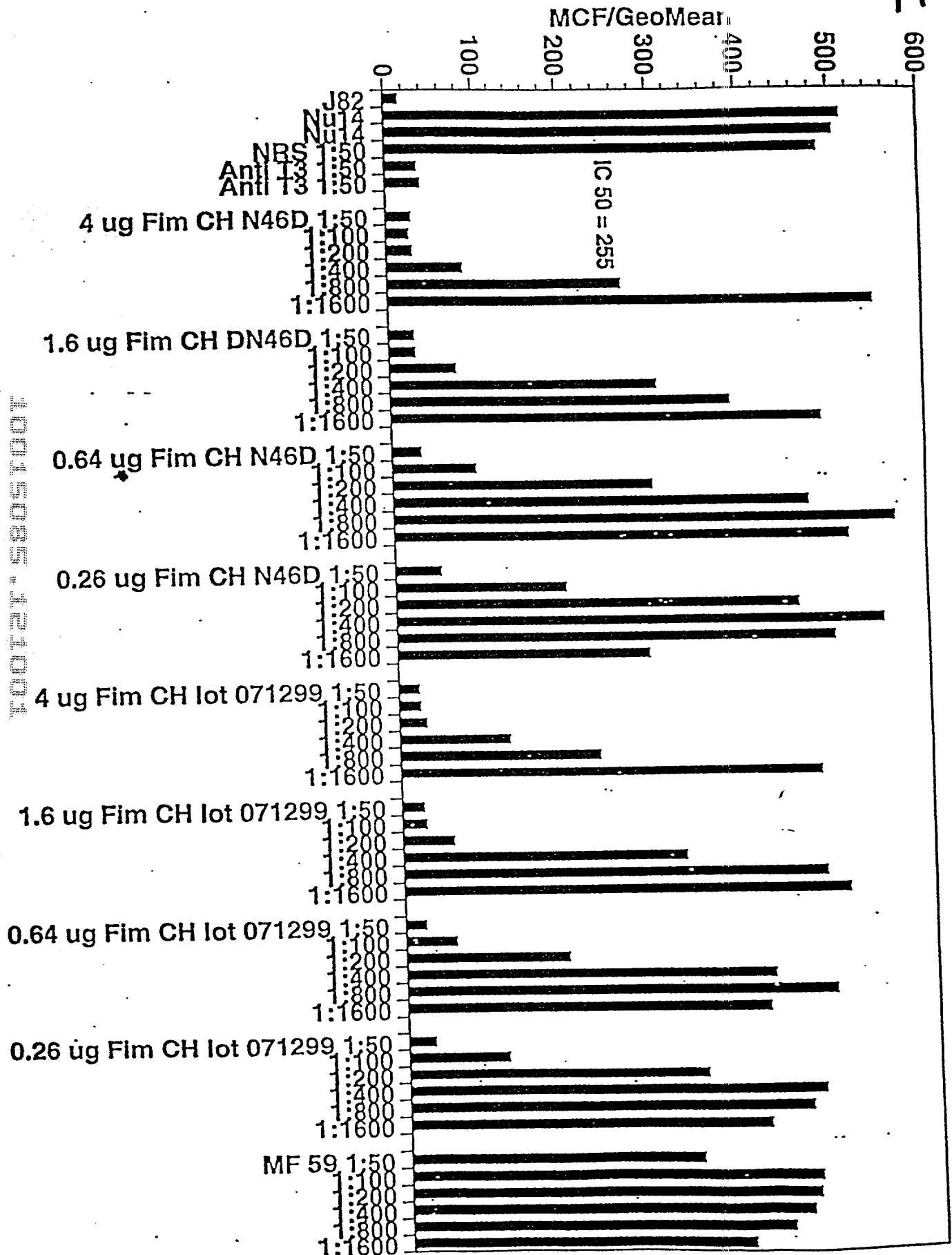


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B

MCF/GeoMear

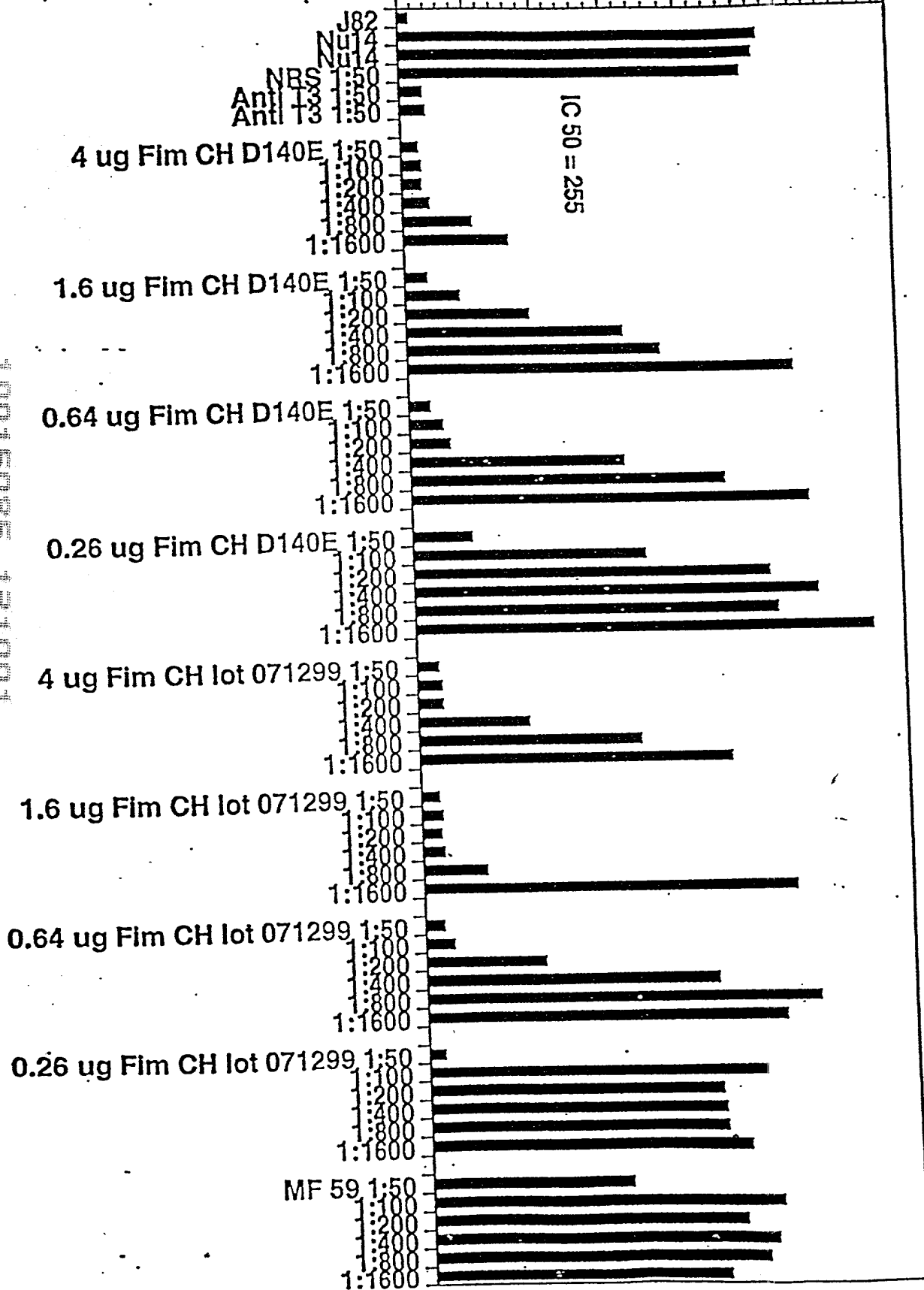


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C

MCF/GeoMean

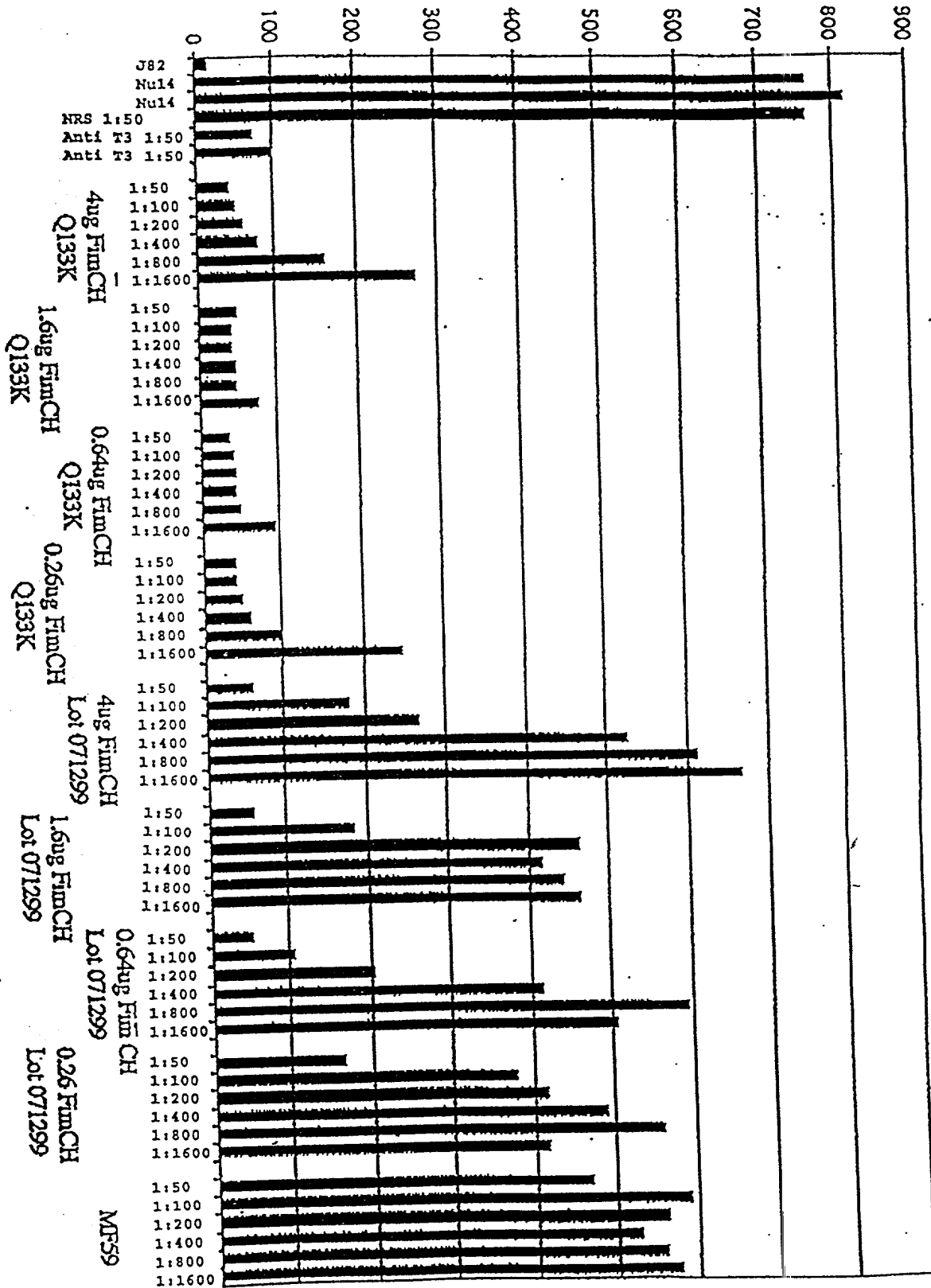


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D

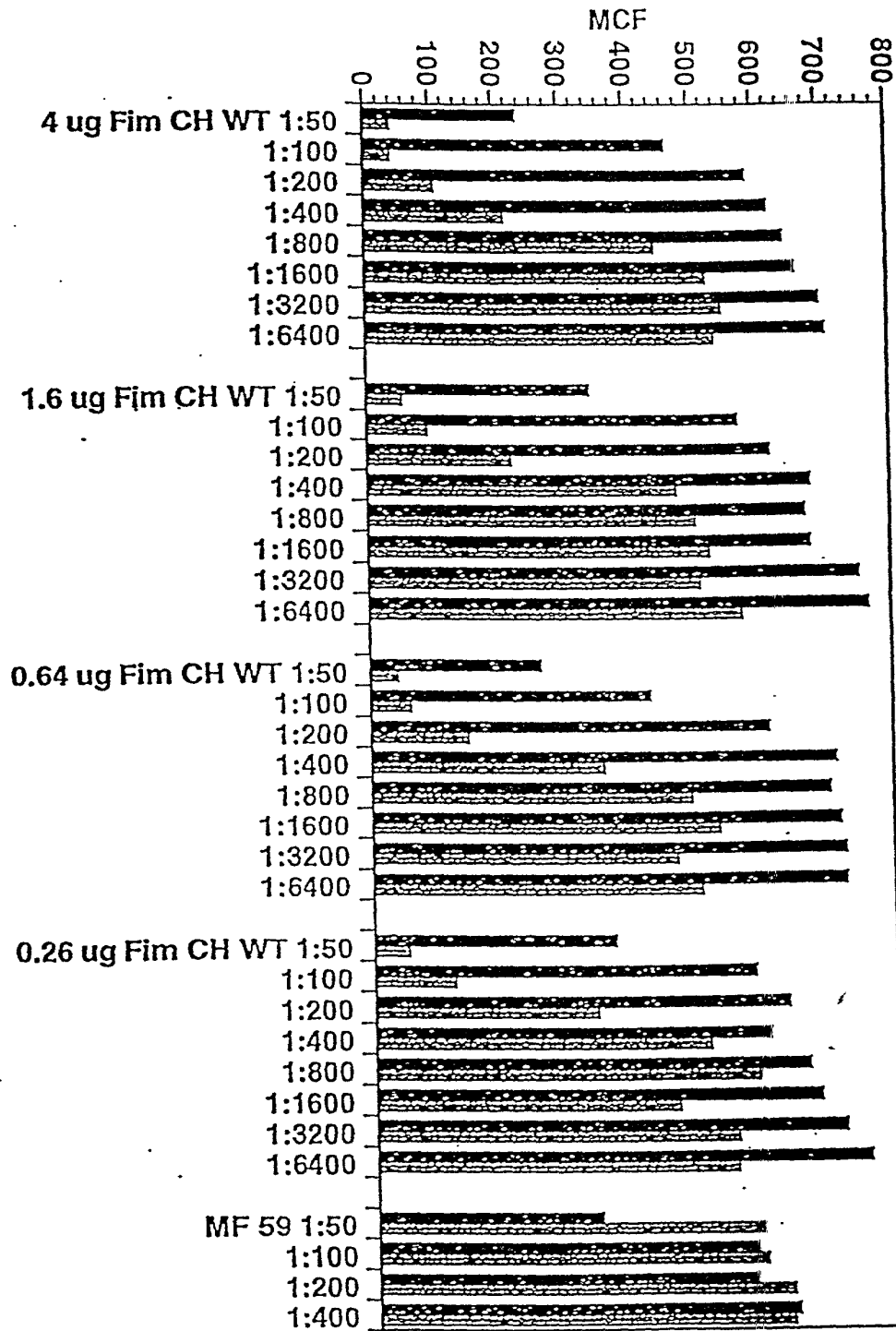


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E

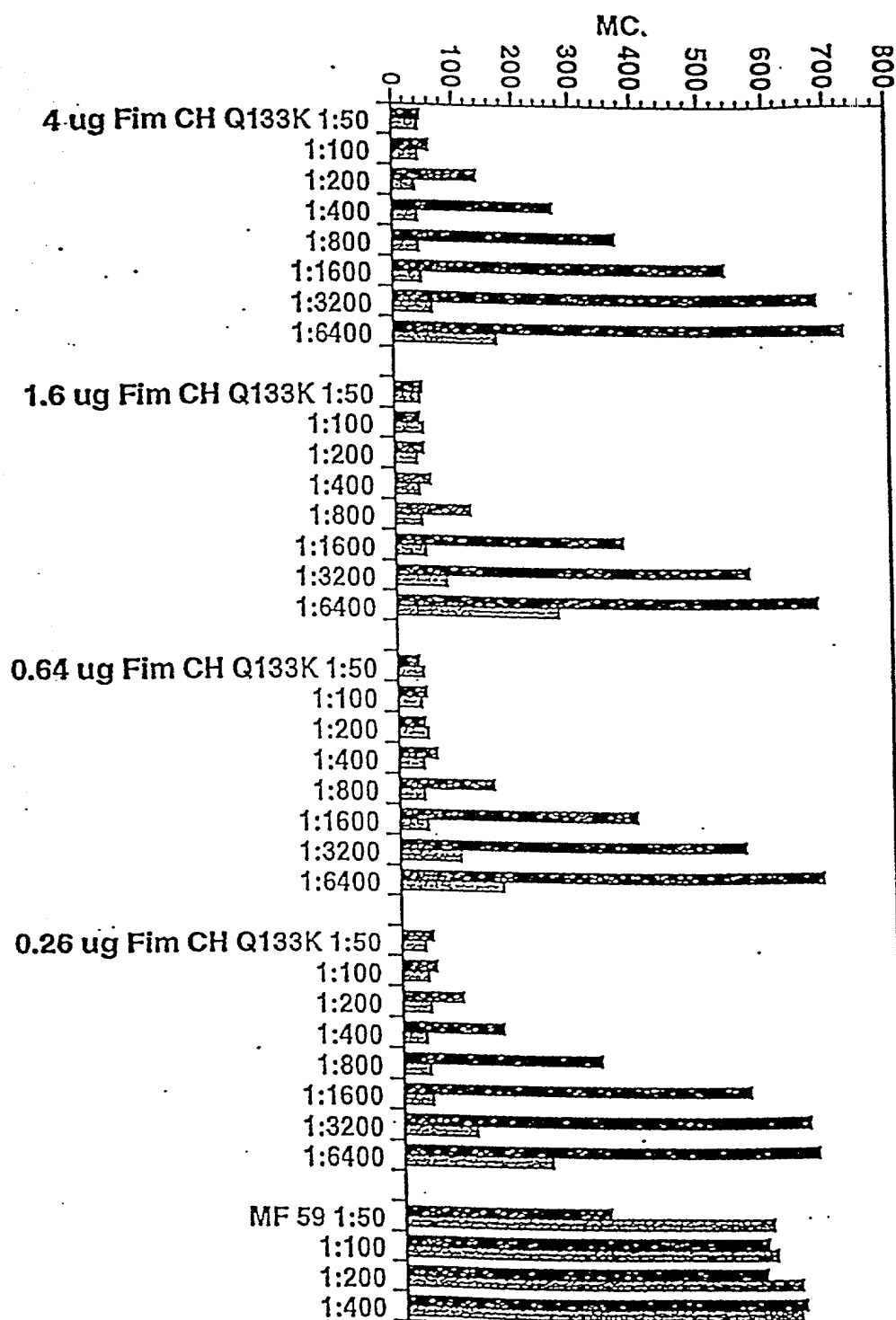


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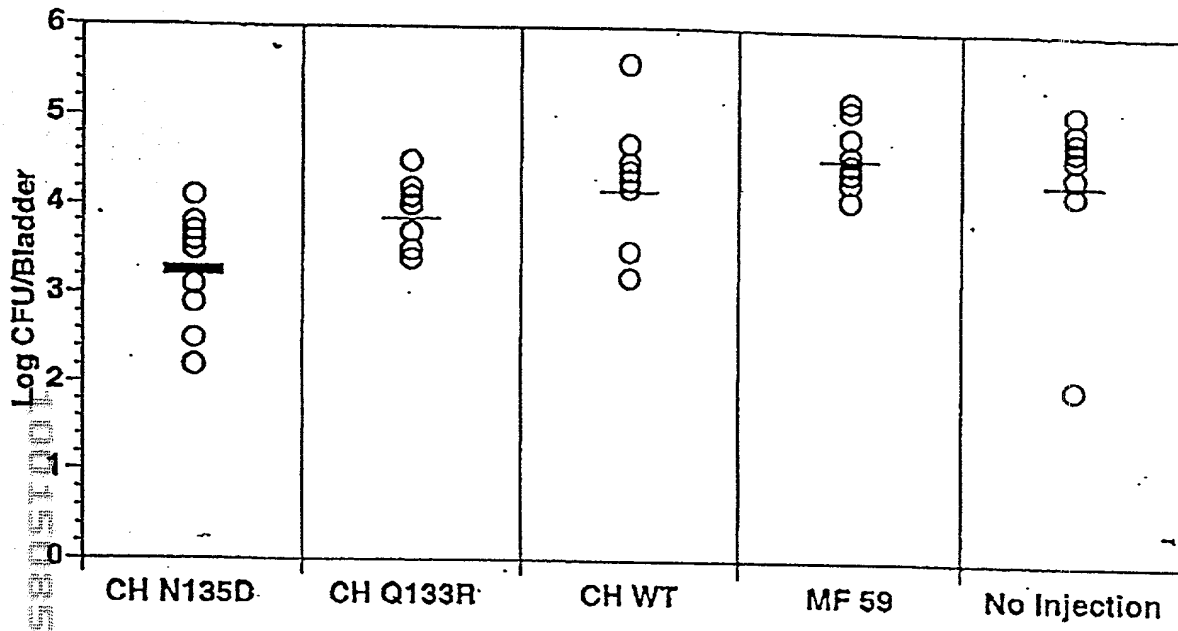


Figure 13
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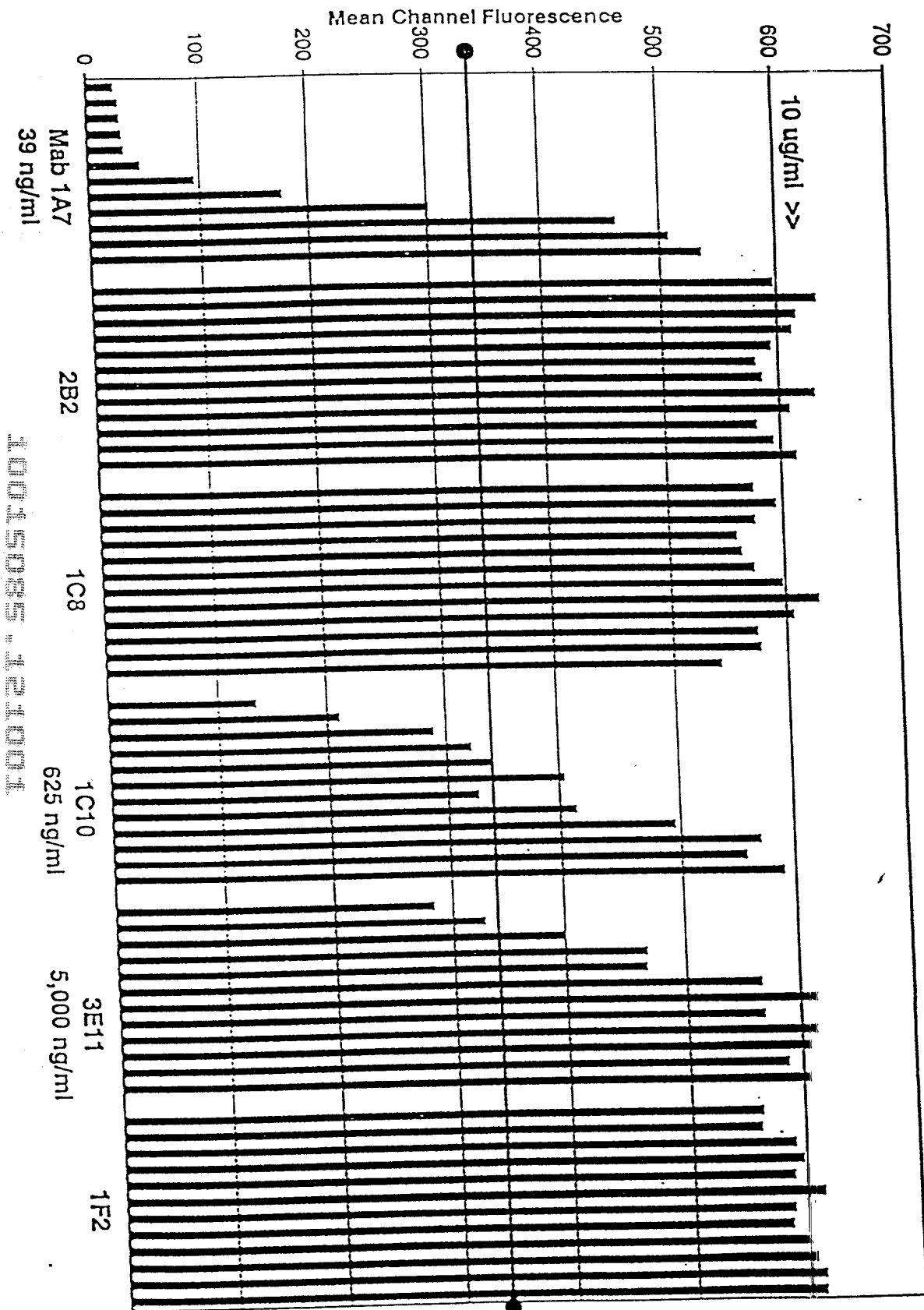


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A

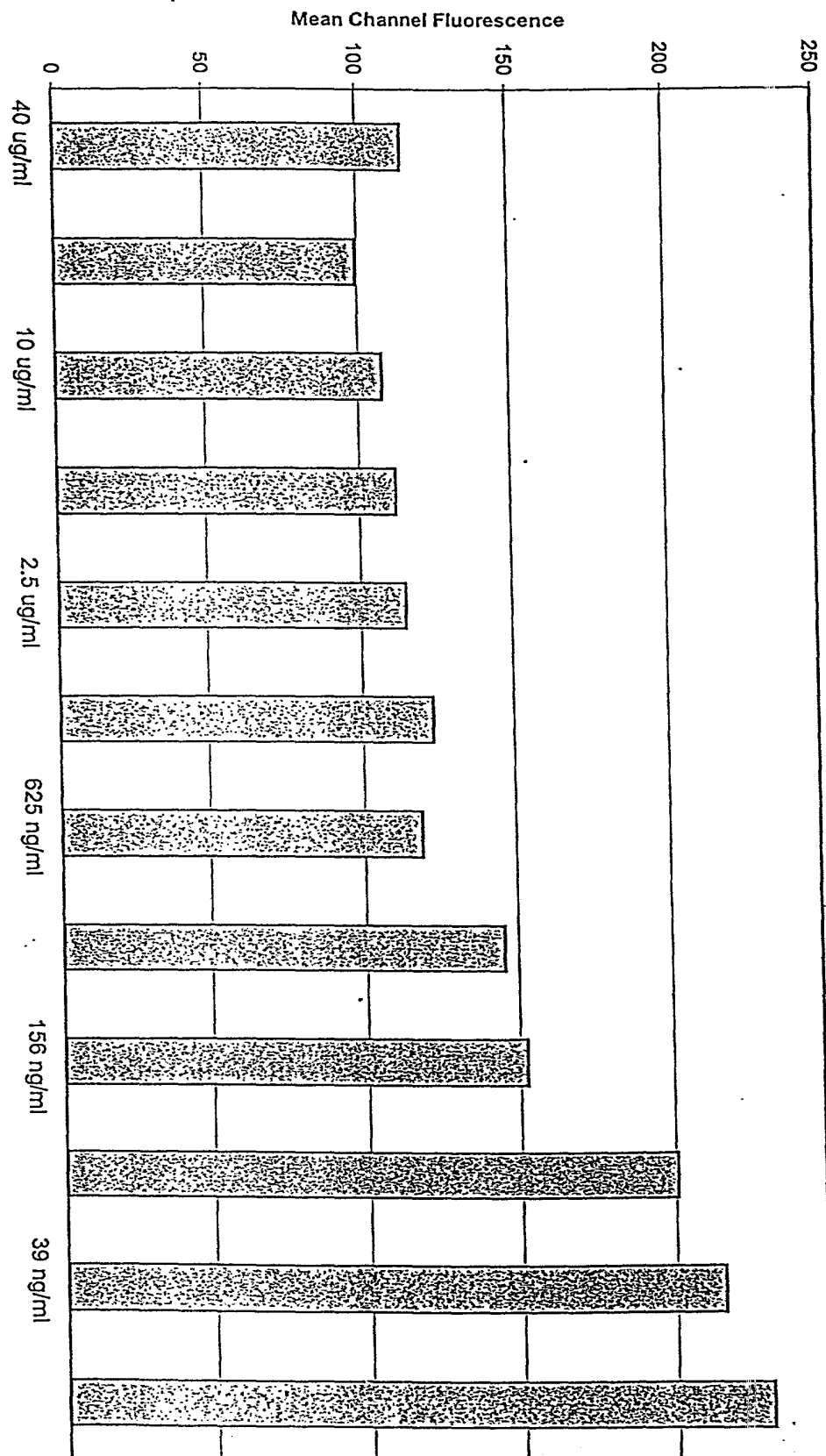


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10015085-131001

B

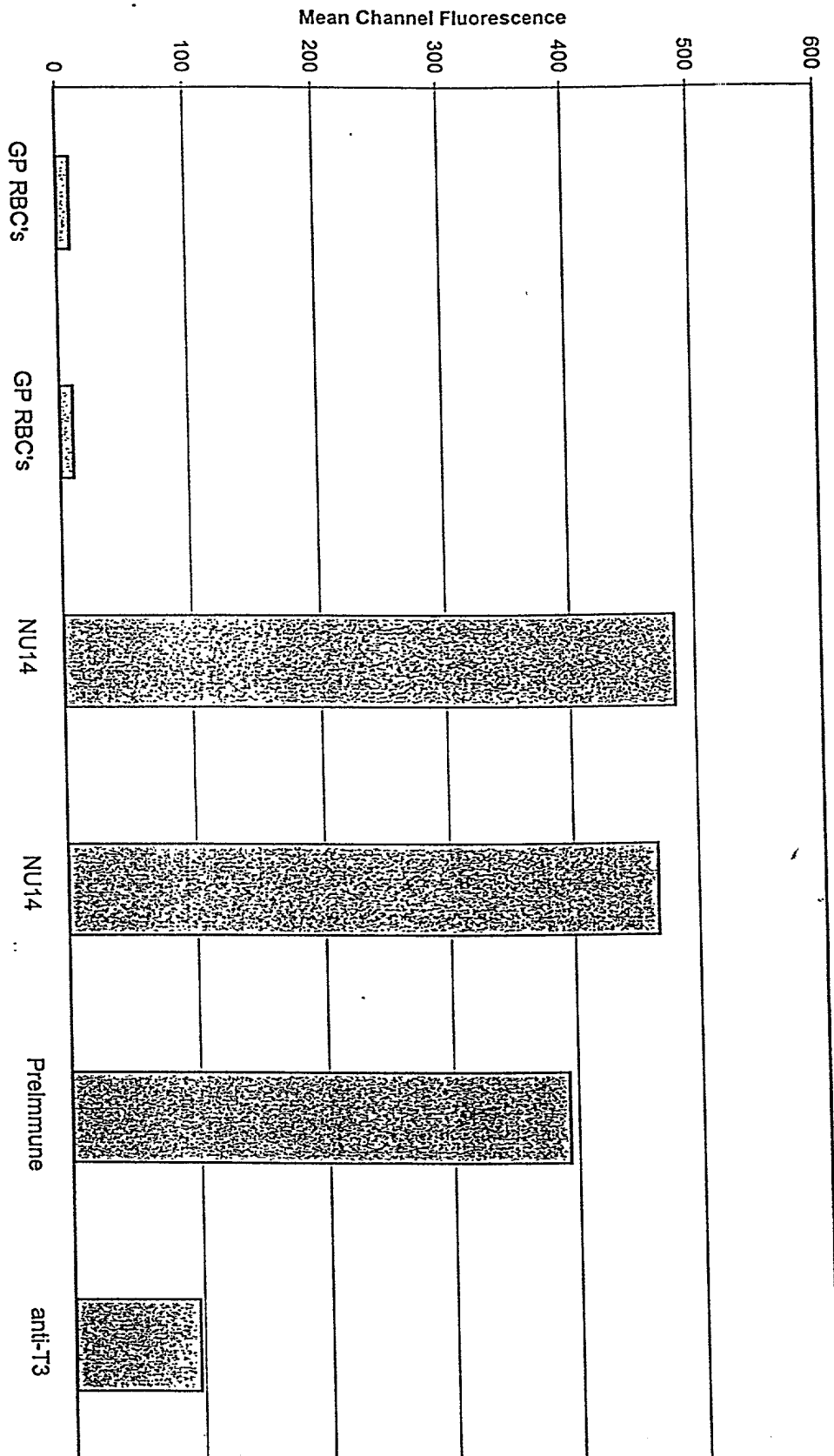


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J0015085 121001

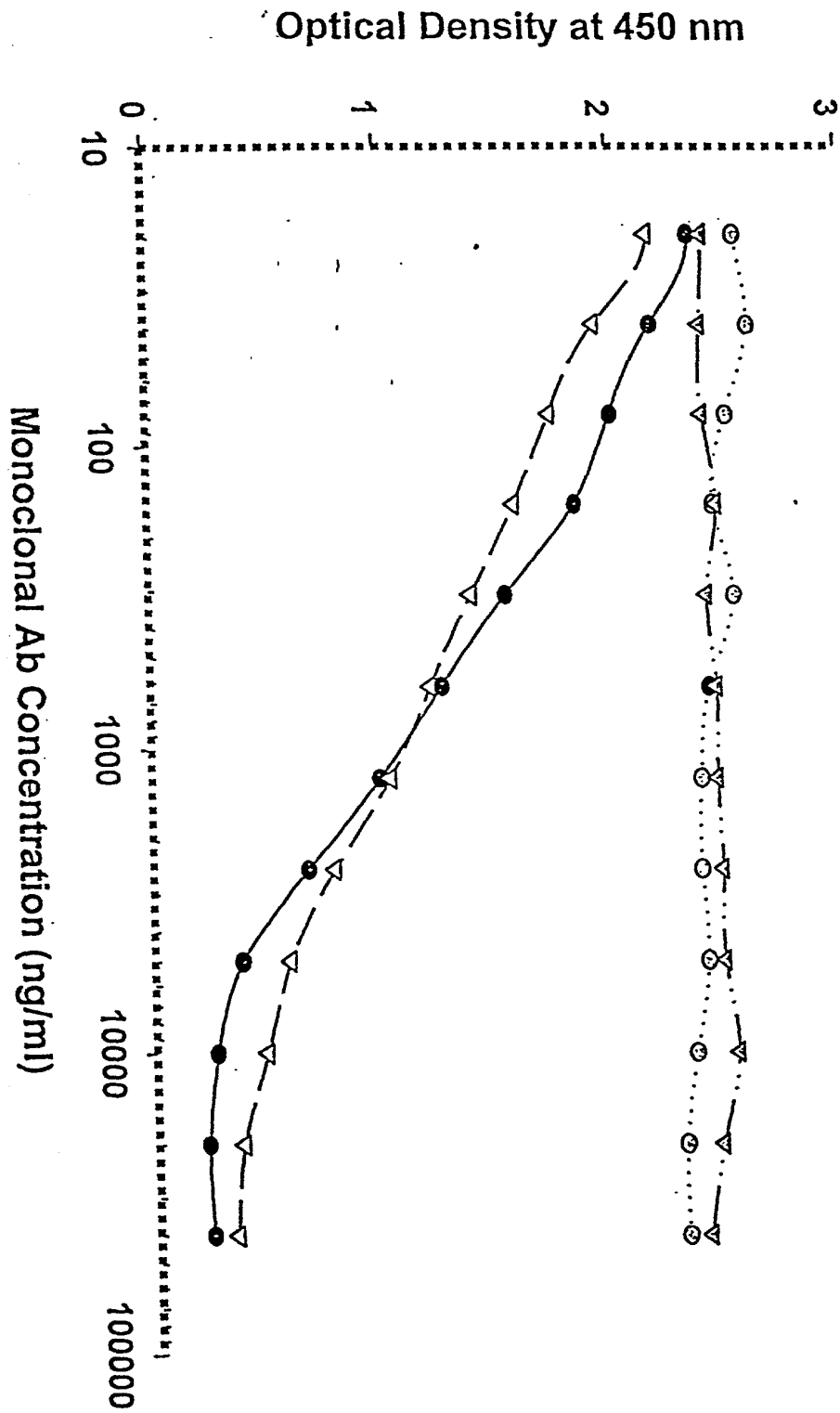


Figure 16
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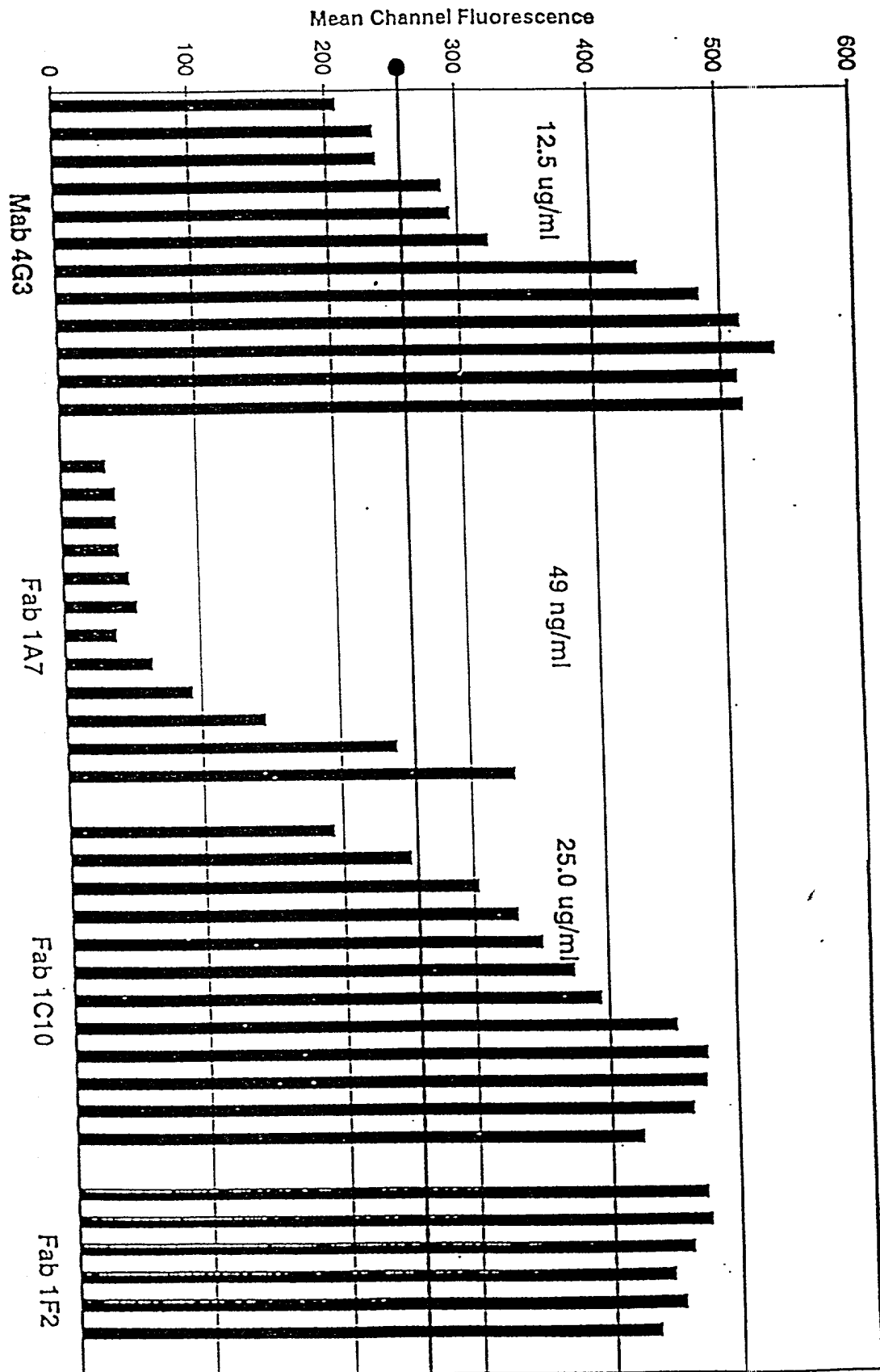


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10015085-121001

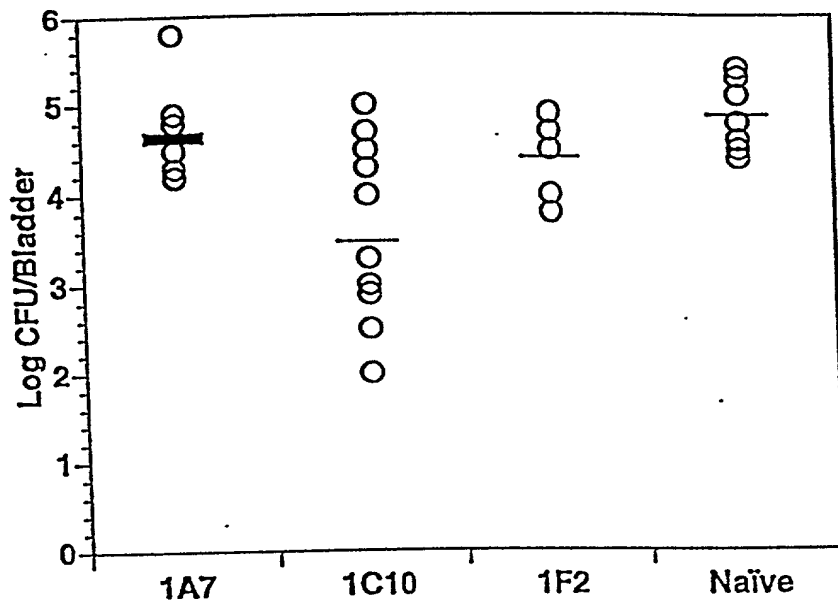
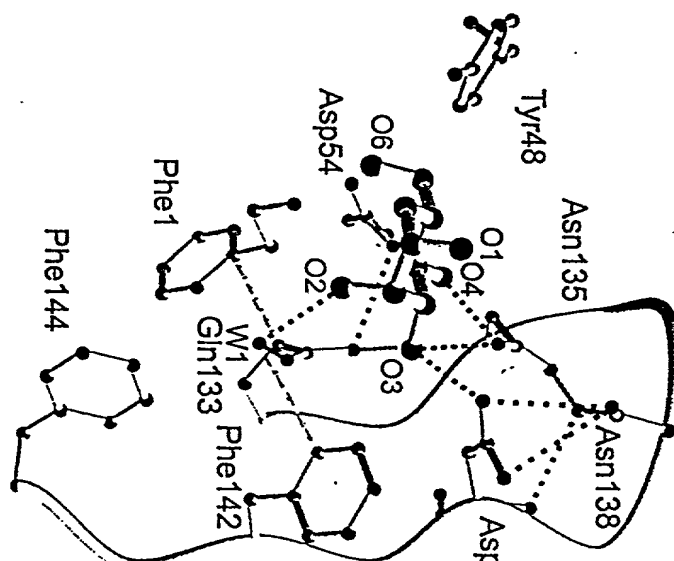


Figure 18
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a



b

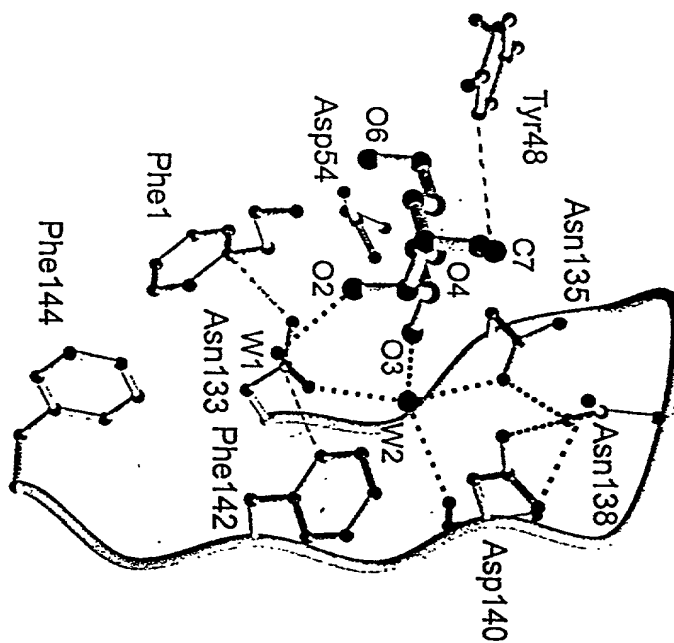


Figure 19
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